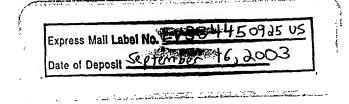
## SEQUENCE LISTING

<110> Altmann, Scott W
Wang, Luquan
Graziano, Michael
Murgolo, Nick



<120> NPC1L1 (NPC3) AND METHODS OF USE THEREOF

<130> JB01603K2 US

<140>

<141> 2003-09-16

<150> 60/397,442

<151> 2002-07-19

<150> 10/621,758

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<151> 2003-08-22

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| atg go<br>Met Al       | ca gct |       |     |   |   |   |   |   |   |   |   |   |   |   | 48  |
| gcg gc<br>Ala Al       |        |       |     |   |   |   |   |   |   |   |   |   |   |   | 96  |
| acc tt<br>Thr Ph       |        |       |     |   |   |   |   |   |   |   |   |   |   |   | 144 |
| acg to<br>Thr Se<br>50 | er Lei |       |     | _ |   | _ | _ |   |   |   | _ | - | _ |   | 192 |
| gtc ac<br>Val Th<br>65 |        | -     |     | _ | - |   |   | _ | - |   | - |   | - | _ | 240 |
| tac aa<br>Tyr As       |        |       |     |   |   |   | - | - | _ |   |   | _ | _ | _ | 288 |
| ctg to<br>Leu Se       |        | _     | _   | - | _ |   |   |   | _ | - |   |   | _ | _ | 336 |
| tgc co<br>Cys Pi       |        | a Cys |     | _ |   |   |   | - |   |   | _ |   |   |   | 384 |
| tgc ag<br>Cys Se<br>13 | -      | _     | _   | _ |   |   |   |   | _ |   |   |   | _ |   | 432 |
| cgg gg<br>Arg G<br>145 |        |       |     |   |   |   |   |   |   |   |   |   |   |   | 480 |
| cag co                 |        |       |     |   |   |   |   |   |   |   |   |   |   |   | 528 |
| atc co                 |        |       | Ala |   |   | - | _ | - |   |   |   |   | _ |   | 576 |
| ggc to<br>Gly Se       |        | a Leu |     |   |   |   |   |   |   |   |   |   |   |   | 624 |
| aca gg                 |        |       | _   | _ | _ | - | _ |   |   |   |   |   |   | _ | 672 |

|   | 210 |   |   |   |   | 215               |   |   |   |   | 220 |   |   |   |   |      |
|---|-----|---|---|---|---|-------------------|---|---|---|---|-----|---|---|---|---|------|
|   |     | _ | _ |   | _ | gat<br>Asp        |   |   | _ |   | _   |   |   | _ |   | 720  |
| _ |     | - |   |   |   | cag<br>Gln        |   | - | - |   | -   | - | _ |   | _ | 768  |
| _ | ~   | _ | - |   |   | tgc<br>Cys        |   | _ |   |   | _   |   |   | _ | _ | 816  |
| - |     |   |   |   | - | ggt<br>Gly        | - | - |   |   |     | _ | - |   |   | 864  |
|   |     |   |   | _ | _ | ttt<br>Phe<br>295 |   | _ |   |   | _   | _ |   |   | - | 912  |
|   | _   |   | _ |   |   | agg<br>Arg        |   | - |   | - | _   | _ |   |   |   | 960  |
| - | -   |   |   |   |   | cat<br>His        | _ |   |   |   |     |   |   |   | _ | 1008 |
| _ |     |   |   |   | _ | aac<br>Asn        |   |   |   |   | -   | _ | _ |   |   | 1056 |
|   |     | _ |   | - | _ | tcc<br>Ser        |   |   | _ |   |     | - |   | _ | _ | 1104 |
|   | _   |   |   |   | - | ctc<br>Leu<br>375 |   |   |   |   | _   | _ | _ |   | _ | 1152 |
|   |     |   |   |   |   | cgg<br>Arg        |   |   |   |   |     |   |   |   |   | 1200 |
|   |     |   |   |   | - | acc<br>Thr        |   | _ |   |   |     |   |   | - |   | 1248 |
|   |     | - |   | _ |   | gac<br>Asp        |   |   | _ |   |     |   |   |   |   | 1296 |
| - |     |   | _ |   | _ | gac<br>Asp        |   | _ | _ |   | _   | _ |   |   | _ | 1344 |
|   |     |   | - |   | _ | caa<br>Gln<br>455 |   |   |   |   |     | _ |   | _ |   | 1392 |

|   |   |   |   | - |   |   |   | gcc<br>Ala        |   |   |   |   |   |   |   | 1440 |
|---|---|---|---|---|---|---|---|-------------------|---|---|---|---|---|---|---|------|
|   |   |   |   | - |   |   |   | agc<br>Ser        |   |   |   |   |   | _ |   | 1488 |
|   | - |   |   | _ | _ |   | - | gcc<br>Ala<br>505 |   | _ |   | _ |   |   | - | 1536 |
|   |   | _ |   | _ |   | - | _ | cat<br>His        |   |   |   | - | _ |   | _ | 1584 |
|   |   | _ |   |   | _ |   | _ | tct<br>Ser        | _ | _ | _ | _ | - | - | _ | 1632 |
|   |   |   |   |   |   |   |   | ttc<br>Phe        |   | - | - |   |   |   |   | 1680 |
|   |   |   |   |   |   |   |   | gcg<br>Ala        |   |   |   |   |   |   |   | 1728 |
|   |   |   |   | - | _ | _ |   | cgc<br>Arg<br>585 | _ | - | - | _ | _ |   |   | 1776 |
|   |   |   |   | _ | - |   | - | gaa<br>Glu        |   |   | _ |   |   |   | _ | 1824 |
| _ | _ |   | _ | - | _ |   |   | gct<br>Ala        |   | _ |   | - | _ | - |   | 1872 |
|   |   |   |   |   |   | ~ | - | ctg<br>Leu        |   | ~ |   | _ | _ | _ |   | 1920 |
|   |   | _ |   |   |   |   |   | ctg<br>Leu        |   |   |   |   |   |   | - | 1968 |
| _ | - | - | - |   |   |   |   | aag<br>Lys<br>665 | - |   | _ |   |   |   |   | 2016 |
|   |   | - |   | - |   | - | - | ctg<br>Leu        | - | _ | _ |   |   |   |   | 2064 |
|   |   |   |   |   |   |   | _ | gtt<br>Val        |   |   |   |   | - |   |   | 2112 |

|     |     |     |     |     |     |     |     | aac<br>Asn        |     |     |     |     |     |     |     | 2160 |
|-----|-----|-----|-----|-----|-----|-----|-----|-------------------|-----|-----|-----|-----|-----|-----|-----|------|
|     |     |     |     |     |     |     |     | Gly<br>ggg        |     |     |     |     |     |     |     | 2208 |
|     |     |     |     |     |     |     |     | ccc<br>Pro<br>745 |     |     |     |     |     |     |     | 2256 |
|     |     |     |     |     |     |     |     | gly<br>ggg        |     |     |     |     |     |     |     | 2304 |
|     |     |     |     | -   | _   |     |     | ggc<br>Gly        |     | _   |     |     |     |     |     | 2352 |
| -   |     | _   | -   |     | -   |     |     | gcc<br>Ala        | _   |     |     | -   | -   | _   | -   | 2400 |
|     | _   |     | _   |     | _   | _   | _   | gtc<br>Val        |     | _   | _   |     |     |     |     | 2448 |
| -   | _   |     |     |     |     | _   |     | gaa<br>Glu<br>825 |     |     |     |     | _   |     |     | 2496 |
| _   | _   |     |     | _   |     |     | _   | ctg<br>Leu        |     | -   |     |     | _   |     | -   | 2544 |
|     | _   | _   | _   |     | _   |     | -   | ttt<br>Phe        |     | _   |     |     |     |     | _   | 2592 |
|     |     |     |     |     |     |     |     | cag<br>Gln        |     | _   |     |     |     |     |     | 2640 |
| _   |     | _   |     | -   |     |     |     | ttt<br>Phe        | -   |     | _   |     |     | -   |     | 2688 |
|     |     |     |     |     |     | _   |     | acc<br>Thr<br>905 | _   |     |     |     |     |     |     | 2736 |
|     | -   |     | _   |     | _   |     | _   | tct<br>Ser        | _   | -   |     | _   | _   | _   |     | 2784 |
|     |     |     | -   |     |     |     |     | gcc<br>Ala        | -   | _   |     |     | -   | _   |     | 2832 |
| tac | gtg | gct | att | gct | gca | tcc | tcc | tgg               | gta | gat | gac | ttc | atc | gac | tgg | 2880 |

| Tyr Val Ala Ile<br>945                    | Ala Ala Ser Se<br>950              | er Trp Val As<br>95 | sp Asp Phe Ile As<br>5                       | 960                     |
|---|------------------------------------|---------------------|--|-------------------------|
| _   | _                                  |                     | at ata cgt ggc cc<br>or Ile Arg Gly Pr<br>97 | o His                   |
|   |                                    |                     | c ttc aac tgc tter<br>Phe Asn Cys Le<br>990  |                         |
| aac tgc atg aac<br>Asn Cys Met Asn<br>995 | Arg Thr Leu G                      |                     |  | gaa cag 3024<br>Glu Gln |
|   |                                    | Phe Leu Asn         | gat ccg ccc aat<br>Asp Pro Pro Asn<br>1020   |                         |
| <b>-</b>                                  | ggg ggt cta<br>Gly Gly Leu<br>1030 | Ala Ala Tyr         | aga acg tct gtg<br>Arg Thr Ser Val<br>1035   |                         |
|   |                                    | Ile Ala Ser         | cag ttc atg gcc<br>Gln Phe Met Ala<br>1050   |                         |
| _   | agg aac tca<br>Arg Asn Ser<br>1060 | Gln Asp Phe         | _  |                         |
|   |                                    | Asn Ile Thr         | gct gac cta cgg<br>Ala Asp Leu Arg<br>1080   | -                       |
|   | gat cca aac<br>Asp Pro Asn<br>1090 | Phe Glu Val         | ttc cct tac acg<br>Phe Pro Tyr Thr<br>1095   |                         |
|   | tac cag caa<br>Tyr Gln Gln<br>1105 | Tyr Leu Thr         | gtc ctt cct gag<br>Val Leu Pro Glu<br>1110   | 1.2                     |
|   |                                    | Phe Val Pro         | acc ttt gtt gtc<br>Thr Phe Val Val<br>1125   |                         |
|   |                                    | Met Cys Ser         | ggg atc ctc aac<br>Gly Ile Leu Asn<br>1140   |                         |
|   |                                    | Val Asp Thr         | att ggc ctc atg<br>Ile Gly Leu Met<br>1155   | _                       |
|   | -                                  | Ala Val Ser         | ctc atc aac ctt<br>Leu Ile Asn Leu<br>1170   | -                       |
|   |                                    |                     | tcc cac atc act<br>Ser His Ile Thr           |                         |

. .

|            | 1175               |            |            |            |            | 1180               |            |            |            |            | 1185               |            |            |            |      |
|------------|--------------------|------------|------------|------------|------------|--------------------|------------|------------|------------|------------|--------------------|------------|------------|------------|------|
|            |                    |            |            |            |            | aag<br>Lys<br>1195 |            |            |            |            |                    |            |            |            | 3609 |
|            | gct<br>Ala<br>1205 |            |            |            |            | ggc<br>Gly<br>1210 |            |            |            |            |                    |            |            |            | 3654 |
|            | acc<br>Thr<br>1220 |            |            |            |            | atc<br>Ile<br>1225 |            |            |            |            |                    |            |            |            | 3699 |
|            |                    |            |            |            |            | ttc<br>Phe<br>1240 |            |            |            |            |                    |            |            |            | 3744 |
| ttg<br>Leu | ctg<br>Leu<br>1250 | ggt<br>Gly | ctg<br>Leu | ctg<br>Leu | cat<br>His | ggc<br>Gly<br>1255 | ctg<br>Leu | gtc<br>Val | ttc<br>Phe | ctg<br>Leu | ccg<br>Pro<br>1260 | gtt<br>Val | gtc<br>Val | ctc<br>Leu | 3789 |
| agc<br>Ser | tat<br>Tyr<br>1265 | ctg<br>Leu | gga<br>Gly | cca<br>Pro | gat<br>Asp | gtt<br>Val<br>1270 | aac<br>Asn | caa<br>Gln | gct<br>Ala | ctg<br>Leu | gta<br>Val<br>1275 | cag<br>Gln | gag<br>Glu | gag<br>Glu | 3834 |
|            |                    |            |            |            |            | gca<br>Ala<br>1285 |            |            |            |            |                    |            | tgc<br>Cys |            | 3879 |
|            |                    |            |            |            |            | gat<br>Asp<br>1300 |            |            |            |            |                    |            | tac<br>Tyr | _          | 3924 |
|            | gcc<br>Ala<br>1310 |            |            |            |            | cac<br>His<br>1315 |            |            |            |            | gct<br>Ala<br>1320 |            |            |            | 3969 |
| _          | ccc<br>Pro<br>1325 | Lys        | _          | _          |            | aag<br>Lys<br>1330 | ttc<br>Phe | taa        |            |            |                    |            |            |            | 3996 |
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| <21        | 1> 1               | 331        |            |            |            |                    |            |            |            |            |                    |            |            |            |      |
| <21        | 2> P               | RT         |            |            |            |                    |            |            |            |            |                    |            |            |            |      |

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<213> Rattus sp.

Met Ala Ala Ala Trp Leu Gly Trp Leu Leu Trp Ala Leu Leu Ser 1 5 10 15

Ala Ala Gl<br/>n Gly Glu Leu Tyr Thr $\mbox{Pro}$  Lys His Glu Ala Gly Val<br/> Cys 20 25 30

Thr Phe Tyr Glu Glu Cys Gly Lys Asn Pro Glu Leu Ser Gly Gly Leu 35 40 45

Thr Ser Leu Ser Asn Val Ser Cys Leu Ser Asn Thr Pro Ala Arg His 50 60

Val Thr Gly Glu His Leu Ala Leu Leu Gln Arg Ile Cys Pro Arg Leu 65 70 75 80

Tyr Asn Gly Pro Asn Thr Thr Phe Ala Cys Cys Ser Thr Lys Gln Leu 85 90 95

Leu Ser Leu Glu Ser Ser Met Ser Ile Thr Lys Ala Leu Leu Thr Arg 100 105 110

Cys Pro Ala Cys Ser Asp Asn Phe Val Ser Leu His Cys His Asn Thr 115 120 125

Cys Ser Pro Asp Gln Ser Leu Phe Ile Asn Val Thr Arg Val Val Glu 130 135 140

Arg Gly Ala Gly Glu Pro Pro Ala Val Val Ala Tyr Glu Ala Phe Tyr 145 150 155 160

Gln Arg Ser Phe Ala Glu Lys Ala Tyr Glu Ser Cys Ser Gln Val Arg 165 170 175

Ile Pro Ala Ala Ala Ser Leu Ala Val Gly Ser Met Cys Gly Val Tyr 180 185 190

Gly Ser Ala Leu Cys Asn Ala Gln Arg Trp Leu Asn Phe Gln Gly Asp 195 200 205

Thr Gly Asn Gly Leu Ala Pro Leu Asp Ile Thr Phe His Leu Leu Glu 210 215 220

Pro Gly Gln Ala Leu Pro Asp Gly Ile Gln Pro Leu Asn Gly Lys Ile 225 230 235 240

Ala Pro Cys Asn Glu Ser Gln Gly Asp Asp Ser Ala Val Cys Ser Cys 245 250 255

Gln Asp Cys Ala Ala Ser Cys Pro Val Ile Pro Pro Pro Glu Ala Leu 260 265 270 Ile Ile Phe Thr Ala Val Phe Val Leu Leu Ser Ala Val Leu Val Arg Leu Arg Val Val Ser Asn Arg Asn Lys Asn Lys Ala Glu Gly Pro Gln Glu Ala Pro Lys Leu Pro His Lys His Lys Leu Ser Pro His Thr Ile Leu Gly Arg Phe Phe Gln Asn Trp Gly Thr Arg Val Ala Ser Trp Pro 340 345 Leu Thr Val Leu Ala Leu Ser Phe Ile Val Val Ile Ala Leu Ala Ala 360 Gly Leu Thr Phe Ile Glu Leu Thr Thr Asp Pro Val Glu Leu Trp Ser 375 380 Ala Pro Lys Ser Gln Ala Arg Lys Glu Lys Ser Phe His Asp Glu His 390 395 Phe Gly Pro Phe Phe Arg Thr Asn Gln Ile Phe Val Thr Ala Arg Asn 405 410 Arg Ser Ser Tyr Lys Tyr Asp Ser Leu Leu Gly Ser Lys Asn Phe 425 Ser Gly Ile Leu Ser Leu Asp Phe Leu Leu Glu Leu Glu Leu Gln 440 Glu Arg Leu Arg His Leu Gln Val Trp Ser Pro Glu Ala Glu Arg Asn Ile Ser Leu Gln Asp Ile Cys Tyr Ala Pro Leu Asn Pro Tyr Asn Thr

Arg Pro Ser Phe Tyr Met Gly Arg Met Pro Gly Trp Leu Ala Leu Ile

Asn Arg Thr Leu Leu Met Leu Thr Ala Asn Gln Thr Leu Asn Gly Gln
500 505 510

Ser Leu Ser Asp Cys Cys Val Asn Ser Leu Leu Gln Tyr Phe Gln Asn

470

485

490

475

Thr Ser Leu Val Asp Trp Lys Asp His Phe Leu Tyr Cys Ala Asn Ala 515 520 525

Pro Leu Thr Phe Lys Asp Gly Thr Ser Leu Ala Leu Ser Cys Met Ala 530

Asp Tyr Gly Ala Pro Val Phe Pro Phe Leu Ala Val Gly Gly Tyr Gln 545 555 555

Gly Thr Asp Tyr Ser Glu Ala Glu Ala Leu Ile Ile Thr Phe Ser Leu 565 570 575

Asn Asn Tyr Pro Ala Asp Asp Pro Arg Met Ala Gln Ala Lys Leu Trp 580 585 590

Glu Glu Ala Phe Leu Lys Glu Met Glu Ser Phe Gln Arg Asn Thr Ser 595 600 605

Asp Lys Phe Gln Val Ala Phe Ser Ala Glu Arg Ser Leu Glu Asp Glu 610 620

Ile Asn Arg Thr Thr Ile Gln Asp Leu Pro Val Phe Ala Val Ser Tyr 625 630 635 640

Ile Ile Val Phe Leu Tyr Ile Ser Leu Ala Leu Gly Ser Tyr Ser Arg 645 650 655

Cys Ser Arg Val Ala Val Glu Ser Lys Ala Thr Leu Gly Leu Gly Gly 660 665 670

Val Ile Val Val Leu Gly Ala Val Leu Ala Ala Met Gly Phe Tyr Ser 675 680 685

Tyr Leu Gly Val Pro Ser Ser Leu Val Ile Ile Gln Val Val Pro Phe 690 695 700

Leu Val Leu Ala Val Gly Ala Asp Asn Ile Phe Ile Phe Val Leu Glu 705 710 715 720

Tyr Gln Arg Leu Pro Arg Met Pro Gly Glu Gln Arg Glu Ala His Ile 725 730 735

Gly Arg Thr Leu Gly Ser Val Ala Pro Ser Met Leu Leu Cys Ser Leu 740 745 750

Ser Glu Ala Ile Cys Phe Phe Leu Gly Ala Leu Thr Pro Met Pro Ala

Val Arg Thr Phe Ala Leu Thr Ser Gly Leu Ala Ile Ile Leu Asp Phe 770 780

765

760

755

- Leu Leu Gln Met Thr Ala Phe Val Ala Leu Leu Ser Leu Asp Ser Lys 785 790 795 800
- Arg Gln Glu Ala Ser Arg Pro Asp Val Leu Cys Cys Phe Ser Thr Arg 805 810 815
- Lys Leu Pro Pro Pro Lys Glu Lys Glu Gly Leu Leu Arg Phe Phe 820 825 830
- Arg Lys Ile Tyr Ala Pro Phe Leu Leu His Arg Phe Ile Arg Pro Val 835 840 845
- Val Met Leu Leu Phe Leu Thr Leu Phe Gly Ala Asn Leu Tyr Leu Met 850 855 860
- Cys Asn Ile Asn Val Gly Leu Asp Gln Glu Leu Ala Leu Pro Lys Asp 865 870 875 880
- Ser Tyr Leu Ile Asp Tyr Phe Leu Phe Leu Asn Arg Tyr Leu Glu Val 885 890 895
- Gly Pro Pro Val Tyr Phe Val Thr Thr Ser Gly Phe Asn Phe Ser Ser 900 910
- Glu Ala Gly Met Asn Ala Thr Cys Ser Ser Ala Gly Cys Lys Ser Phe 915 920 925
- Ser Leu Thr Gln Lys Ile Gln Tyr Ala Ser Glu Phe Pro Asp Gln Ser 930 935 940
- Tyr Val Ala Ile Ala Ala Ser Ser Trp Val Asp Asp Phe Ile Asp Trp 945 950 955 960
- Leu Thr Pro Ser Ser Ser Cys Cys Arg Leu Tyr Ile Arg Gly Pro His 965 970 975
- Lys Asp Glu Phe Cys Pro Ser Thr Asp Thr Ser Phe Asn Cys Leu Lys 980 985 990

Phe His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Pro Pro Asn Ile Arg Cys Pro Lys Gly Gly Leu Ala Ala Tyr Arg Thr Ser Val Asn Leu Ser Ser Asp Gly Gln Val Ile Ala Ser Gln Phe Met Ala Tyr His Lys Pro Leu Arg Asn Ser Gln Asp Phe Thr Glu Ala Leu Arg Ala Ser Arg Leu Leu Ala Ala Asn Ile Thr Ala Asp Leu Arg Lys Val Pro Gly Thr Asp Pro Asn Phe Glu Val Phe Pro Tyr Thr Ile Ser Asn Val Phe Tyr Gln Gln Tyr Leu Thr Val Leu Pro Glu Gly Ile Phe Thr Leu Ala Leu Cys Phe Val Pro Thr Phe Val Val Cys Tyr Leu Leu Gly Leu Asp Met Cys Ser Gly Ile Leu Asn Leu Leu Ser Ile Ile Met Ile Leu Val Asp Thr Ile Gly Leu Met Ala Val Trp Gly Ile Ser Tyr Asn Ala Val Ser Leu Ile Asn Leu Val Thr Ala Val Gly Met Ser Val Glu Phe Val Ser His Ile Thr Arg Ser Phe Ala Val Ser Thr Lys Pro Thr Arg Leu Glu Arg Ala Lys Asp Ala Thr Val Phe Met Gly Ser Ala Val Phe Ala Gly Val Ala Met Thr Asn Phe Pro Gly Ile Leu Ile Leu Gly Phe Ala Gln Ala 1220 . 1225 

| Gln Leu Ile Gln Ile Phe Phe Phe Arg Leu Asn Leu Leu Ile Thr<br>1235 1240 1245   |
|---|
| Leu Leu Gly Leu His Gly Leu Val Phe Leu Pro Val Val Leu<br>1250 1255 1260   |
| Ser Tyr Leu Gly Pro Asp Val Asn Gln Ala Leu Val Gln Glu Glu<br>1265 1270 1275   |
| Lys Leu Ala Ser Glu Ala Ala Val Ala Pro Glu Pro Ser Cys Pro<br>1280 1285 1290   |
| Gln Tyr Pro Ser Pro Ala Asp Ala Asp Ala Asn Val Asn Tyr Gly 1295 1300 1305  |
| Phe Ala Pro Glu Leu Ala His Gly Ala Asn Ala Ala Arg Ser Ser<br>1310 1315 1320   |
| Leu Pro Lys Ser Asp Gln Lys Phe<br>1325 1330  |
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| <213> Homo sapiens  |
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| <400> 3 atg gcg gag gcc ctg agg ggc tgg ctg ctg tgg gcc ctg ctc ctg 48  |
| Met Ala Glu Ala Gly Leu Arg Gly Trp Leu Leu Trp Ala Leu Leu  1 5 10 15  |
| cgc ttg gcc cag agt gag cct tac aca acc atc cac cag cct ggc tac 96 Arg Leu Ala Gln Ser Glu Pro Tyr Thr Thr Ile His Gln Pro Gly Tyr 20 25 30 |
| tgc gcc ttc tat gac gaa tgt ggg aag aac cca gag ctg tct gga agc  Cys Ala Phe Tyr Asp Glu Cys Gly Lys Asn Pro Glu Leu Ser Gly Ser  35 40 45  |

| ctc<br>Leu         | atg<br>Met<br>50  | aca<br>Thr        | ctc<br>Leu        | tcc<br>Ser        | aac<br>Asn        | gtg<br>Val<br>55  | tcc<br>Ser        | tgc<br>Cys        | ctg<br>Leu        | tcc<br>Ser        | aac<br>Asn<br>60  | acg<br>Thr        | ccg<br>Pro        | gcc<br>Ala        | cgc<br>Arg        | 192 |
|--------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-----|
| aag<br>Lys<br>65   | atc<br>Ile        | aca<br>Thr        | ggt<br>Gly        | gat<br>Asp        | cac<br>His<br>70  | ctg<br>Leu        | atc<br>Ile        | cta<br>Leu        | tta<br>Leu        | cag<br>Gln<br>75  | aag<br>Lys        | atc<br>Ile        | tgc<br>Cys        | ccc<br>Pro        | cgc<br>Arg<br>80  | 240 |
| ctc<br>Leu         | tac<br>Tyr        | acc<br>Thr        | ggc<br>Gly        | ccc<br>Pro<br>85  | aac<br>Asn        | acc<br>Thr        | caa<br>Gln        | gcc<br>Ala        | tgc<br>Cys<br>90  | tgc<br>Cys        | tcc<br>Ser        | gcc<br>Ala        | aag<br>Lys        | cag<br>Gln<br>95  | ctg<br>Leu        | 288 |
| gta<br>Val         | tca<br>Ser        | ctg<br>Leu        | gaa<br>Glu<br>100 | gcg<br>Ala        | agt<br>Ser        | ctg<br>Leu        | tcg<br>Ser        | atc<br>Ile<br>105 | acc<br>Thr        | aag<br>Lys        | gcc<br>Ala        | ctc<br>Leu        | ctc<br>Leu<br>110 | acc<br>Thr        | cgc<br>Arg        | 336 |
| tgc<br>Cys         | cca<br>Pro        | gcc<br>Ala<br>115 | tgc<br>Cys        | tct<br>Ser        | gac<br>Asp        | aat<br>Asn        | ttt<br>Phe<br>120 | gtg<br>Val        | aac<br>Asn        | ctg<br>Leu        | cac<br>His        | tgc<br>Cys<br>125 | cac<br>His        | aac<br>Asn        | acg<br>Thr        | 384 |
| tgc<br>Cys         | agc<br>Ser<br>130 | ccc<br>Pro        | aat<br>Asn        | cag<br>Gln        | agc<br>Ser        | ctc<br>Leu<br>135 | ttc<br>Phe        | atc<br>Ile        | aat<br>Asn        | gtg<br>Val        | acc<br>Thr<br>140 | cgc<br>Arg        | gtg<br>Val        | gcc<br>Ala        | cag<br>Gln        | 432 |
| cta<br>Leu<br>145  | Gly               | gct<br>Ala        | gga<br>Gly        | caa<br>Gln        | ctc<br>Leu<br>150 | cca<br>Pro        | gct<br>Ala        | gtg<br>Val        | gtg<br>Val        | gcc<br>Ala<br>155 | tat<br>Tyr        | gag<br>Glu        | gcc<br>Ala        | ttc<br>Phe        | tac<br>Tyr<br>160 | 480 |
| cag<br>Gln         | cat<br>His        | agc<br>Ser        | ttt<br>Phe        | gcc<br>Ala<br>165 | gag<br>Glu        | cag<br>Gln        | agc<br>Ser        | tat<br>Tyr        | gac<br>Asp<br>170 | tcc<br>Ser        | tgc<br>Cys        | agc<br>Ser        | cgt<br>Arg        | gtg<br>Val<br>175 | cgc<br>Arg        | 528 |
| gtc<br>Val         | cct<br>Pro        | gca<br>Ala        | gct<br>Ala<br>180 | gcc<br>Ala        | acg<br>Thr        | ctg<br>Leu        | gct<br>Ala        | gtg<br>Val<br>185 | Gly               | acc<br>Thr        | atg<br>Met        | tgt<br>Cys        | ggc<br>Gly<br>190 | gtg<br>Val        | tat<br>Tyr        | 576 |
| ggc<br>Gly         | tct<br>Ser        | gcc<br>Ala<br>195 | Leu               | tgc<br>Cys        | aat<br>Asn        | gcc<br>Ala        | cag<br>Gln<br>200 | cgc<br>Arg        | tgg<br>Trp        | ctc<br>Leu        | aac<br>Asn        | ttc<br>Phe<br>205 | cag<br>Gln        | gga<br>Gly        | gac<br>Asp        | 624 |
| aca<br>Thr         | ggc<br>Gly<br>210 | Asn               | ggt<br>Gly        | ctg<br>Leu        | gcc<br>Ala        | cca<br>Pro<br>215 | Leu               | gac<br>Asp        | atc<br>Ile        | acc<br>Thr        | ttc<br>Phe<br>220 | cac<br>His        | ctc<br>Leu        | ttg<br>Leu        | gag<br>Glu        | 672 |
| cct<br>Pro<br>225  | Gly               | cag<br>Gln        | gcc<br>Ala        | gtg<br>Val        | ggg<br>Gly<br>230 | Ser               | ggg<br>Gly        | att<br>Ile        | cag<br>Gln        | cct<br>Pro<br>235 | Leu               | aat<br>Asn        | gag<br>Glu        | ggg               | gtt<br>Val<br>240 | 720 |
| gca<br>Ala         | cgt<br>Arg        | tgc<br>Cys        | aat<br>Asn        | gag<br>Glu<br>245 | Ser               | caa<br>Gln        | ggt<br>Gly        | gac<br>Asp        | gac<br>Asp<br>250 | Val               | gcg<br>Ala        | acc<br>Thr        | tgc<br>Cys        | Ser<br>255        |                   | 768 |
| caa<br>Glr         | a gac<br>n Asp    | tgt<br>Cys        | gct<br>Ala<br>260 | Ala               | tcc<br>Ser        | tgt<br>Cys        | cct               | gcc<br>Ala<br>265 | Ile               | gcc<br>Ala        | cgc<br>Arg        | ccc<br>Pro        | cag<br>Gln<br>270 | Ala               | ctc               | 816 |
| gao<br>As <u>r</u> | c tcc<br>Ser      | acc<br>Thi        | : Phe             | tac<br>Tyr        | ctg<br>Lev        | ggc<br>Gly        | cac<br>Glr<br>280 | n Met             | ccg<br>Pro        | ggc<br>Gly        | agt<br>Ser        | ctg<br>Leu<br>285 | ı Val             | cto<br>Leu        | atc<br>Ile        | 864 |

| atc a             | atc<br>Ile<br>290  | ctc<br>Leu       | tgc<br>Cys         | tc<br>Se                  | t gt<br>r Va      | at F                      | tc<br>he           | gct<br>Ala         | gtg<br>Val         | gt<br>Va       | с а<br>1 Т        | . 111             | atc<br>Ile<br>300 | ct.<br>Le          | g c<br>u L         | tt (<br>eu V           | gtg<br>Val         | gg<br>Gl       | а<br>У            | 912  |
|-------------------|--------------------|------------------|--------------------|---------------------------|-------------------|---------------------------|--------------------|--------------------|--------------------|----------------|-------------------|-------------------|-------------------|--------------------|--------------------|------------------------|--------------------|----------------|-------------------|------|
| ttc<br>Phe<br>305 | cgt<br>Arg         | gtg<br>Val       | gco<br>Ala         | c cc<br>a Pr              | O A.              | cc a<br>la <i>P</i><br>10 | agg<br>Arg         | gac<br>Asp         | aaa<br>Lys         | ag<br>Se       | ; r r             | aag<br>Lys<br>315 | atg<br>Met        | gt<br>Va           | g g                | ac<br>sp               | ccc<br>Pro         | аа<br>Ьу<br>32 |                   | 960  |
| aag<br>Lys        | ggc<br>Gly         | acc<br>Thr       | ago<br>Se:         | c ct<br>r Le<br>32        | eu S              | ct (<br>er <i>i</i>       | gac<br>Asp         | aag<br>Lys         | ctc<br>Leu         | ag<br>Se<br>33 | ar i              | ttc<br>Phe        | tcc<br>Ser        | ac<br>Th           | c c<br>ir F        |                        | acc<br>Thr<br>335  | ct<br>Le       | c<br>eu           | 1008 |
| ctt<br>Leu        | ggc<br>Gly         | caç<br>Glr       | tt<br>Ph<br>34     | e Ph                      | c c<br>ne G       | ag<br>ln                  | ggc<br>Gly         | tgg<br>Trp         | ggc<br>Gly<br>345  | 11             | eg<br>nr '        | tgg<br>Trp        | gtg<br>Val        | gg<br>Al           |                    | ccg<br>Ser<br>350      | tgg<br>Trp         | P              | ct<br>ro          | 1056 |
| ctg<br>Leu        | acc<br>Thr         | ato<br>I10<br>35 | e Le               | g gt<br>u Va              | tg c              | ta<br>eu                  | tct<br>Ser         | gtc<br>Val<br>360  | TTE                | : co           | cg<br>ro          | gtg<br>Val        | gtg<br>Va]        | L A.               | cc 1<br>la 1<br>65 | ttg<br>Leu             | gca<br>Ala         | g<br>A         | cg<br>la          | 1104 |
| ggc<br>Gly        | ctg<br>Leu<br>370  | . Va             | c tt<br>l Ph       | t a                       | ca g<br>hr (      | gaa<br>Slu                | ctc<br>Leu<br>375  | act<br>Thr         | acç<br>Thi         | g g            | ac<br>sp          | ccc<br>Pro        | gtg<br>Va:<br>380 |                    | ag<br>lu           | ctg<br>Leu             | tgg<br>Trp         | t<br>S         | cg<br>er          | 1152 |
| gcc<br>Ala<br>385 | Pro                | aa<br>As         | c aç<br>n Se       | jc c<br>er G              | ln A              | gcc<br>Ala<br>390         | cgg<br>Arg         | agt<br>Sei         | gaç<br>Glı         | g a<br>ı L     | aa<br>ys          | gct<br>Ala<br>395 | 1 11              | сс<br>e H          | at<br>is           | gac<br>Asp             | cag<br>Glr         |                | at<br>is<br>00    | 1200 |
| ttc<br>Phe        | ggo<br>Gly         | c cc<br>y Pr     | c ti               | ne F                      | tc<br>he          | cga<br>Arg                | acc<br>Thr         | aac<br>Ası         | c ca<br>n Gl:      | n v            | gtg<br>Val        | ato               | ct<br>Le          | g a<br>u T         | icg<br>hr          | gct<br>Ala             | Pro<br>415         | -              | ac<br>Asn         | 1248 |
| cgg               | g tc<br>g Se:      | c aç<br>r Se     | er T               | ac a<br>yr <i>P</i><br>20 | agg<br>Arg        | tat<br>Tyr                | gac<br>As <u>r</u> | tc<br>Se           | t ct<br>r Le<br>42 | u i            | ctg<br>Leu        | ctç<br>Lei        | g gg<br>1 Gl      | g c                | ccc<br>Pro         | aag<br>Lys<br>430      |                    | c t            | tc<br>Phe         | 1296 |
| ago<br>Sei        | c gg<br>c Gl       | y I              | cc c<br>le L<br>35 | tg (<br>eu <i>l</i>       | gac<br>Asp        | ctg<br>Leu                | ga<br>As           | c tt<br>o Le<br>44 | u Le               | g o            | ctg<br>Leu        | gaq<br>Glu        | g ct<br>ı L∈      | - u 1              | cta<br>Leu<br>445  | gag                    | ct.<br>Le          | g (<br>u (     | cag<br>Gln        | 1344 |
| ga<br>Gl          | g ag<br>u Ar<br>45 | g L              | tg c<br>eu A       | gg (                      | cac<br>His        | ctc<br>Leu                | ca<br>Gl<br>45     | g gt<br>n Va<br>5  | a to<br>1 Ti       | .b<br>ia       | tcg<br>Ser        | r CC              | o G.              | aa g<br>lu 2<br>60 | gca<br>Ala         | caç<br>Glr             | g cg<br>n Ar       | c<br>g         | aac<br>Asn        | 1392 |
| at<br>Il<br>46    | e Se               | c c<br>er L      | tg c               | ag<br>Sln                 | gac<br>Asp        | atc<br>11e<br>470         | э Су               | c ta               | ıc go<br>r A.      | cc<br>la       | ccc               | c ct<br>Le<br>47  | u A               | at<br>sn           | ccg<br>Pro         | gao<br>As <sub>l</sub> | c aa<br>o As       | t<br>n         | acc<br>Thr<br>480 | 1440 |
| ag<br>Se          | t ct<br>r Le       | cc t<br>eu T     | ac (               | gac<br>Asp                | tgc<br>Cys<br>485 | tgo<br>Cy:                | c at               | .c aa<br>.e As     | ac a<br>sn S       | gc<br>er       | cto<br>Lev<br>490 | и пе              | g c<br>u G        | ag<br>ln           | tat<br>Tyr         | tt.<br>Ph              | c ca<br>e Gl<br>49 |                | aac<br>Asn        | 1488 |
| aa<br>As          | ac ce              | gc a<br>rg T     | hr                 | ctc<br>Leu<br>500         | ctg<br>Leu        | ct                        | g ct<br>u Le       | c a<br>eu T        | nr A               | сс<br>1а<br>05 | aa<br>Asi         | c ca<br>n Gl      | ıg a<br>.n T      | ca<br>hr           | cto                | g at<br>u Me<br>51     |                    | JÀ<br>33       | cag<br>Gln        | 1536 |
| ac<br>Tl          | cc t<br>nr S       | er (             | caa<br>Gln<br>515  | gtc<br>Val                | gac<br>Asp        | tg<br>Tr                  | g aa<br>p Ly       | ys A               | ac c<br>sp H<br>20 | at             | tt<br>Ph          | t ct<br>e Le      | g t<br>eu T       | ac<br>Yr           | tgt<br>Cys<br>52!  | , ,,,                  | c a                | at<br>sn       | gcc<br>Ala        | 1584 |
| C                 | cg c               | tc               | acc                | ttc                       | aag               | g ga                      | t g                | gc a               | ca g               | icc            | ct                | g g               | ec c              | etg                | ag                 | c tç                   | gc a               | tg             | gct               | 1632 |

| Pro               | Leu<br>530        | Thr               | Phe               | Lys               | Asp               | Gly<br>535        | Thr               | Ala               | Leu               | Ala               | Leu<br>540        |                   | Cys               | Met               | Ala               |      |
|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|------|
| gac<br>Asp<br>545 | Tyr               | Gly               | gcc<br>Ala        | cct<br>Pro        | gtc<br>Val<br>550 | ttc<br>Phe        | ccc<br>Pro        | ttc<br>Phe        | ctt<br>Leu        | gcc<br>Ala<br>555 | att<br>Ile        | gly               | Gly               | tac<br>Tyr        | aaa<br>Lys<br>560 | 1680 |
| gga<br>Gly        | aag<br>Lys        | gac<br>Asp        | tat<br>Tyr        | tct<br>Ser<br>565 | Glu               | gca<br>Ala        | gag<br>Glu        | gcc<br>Ala        | ctg<br>Leu<br>570 | atc<br>Ile        | atg<br>Met        | acg<br>Thr        | ttc<br>Phe        | tcc<br>Ser<br>575 | ctc<br>Leu        | 1728 |
| aac<br>Asn        | aat<br>Asn        | tac<br>Tyr        | cct<br>Pro<br>580 | gcc<br>Ala        | Gly<br>aaa        | gac<br>Asp        | ccc<br>Pro        | cgt<br>Arg<br>585 | ctg<br>Leu        | gcc<br>Ala        | cag<br>Gln        | gcc<br>Ala        | aag<br>Lys<br>590 | ctg<br>Leu        | tgg<br>Trp        | 1776 |
| gag<br>Glu        | gag<br>Glu        | gcc<br>Ala<br>595 | ttc<br>Phe        | tta<br>Leu        | gag<br>Glu        | gaa<br>Glu        | atg<br>Met<br>600 | cga<br>Arg        | gcc<br>Ala        | ttc<br>Phe        | cag<br>Gln        | cgt<br>Arg<br>605 | cgg<br>Arg        | atg<br>Met        | gct<br>Ala        | 1824 |
| ggc               | atg<br>Met<br>610 | ttc<br>Phe        | cag<br>Gln        | gtc<br>Val        | acg<br>Thr        | ttc<br>Phe<br>615 | acg<br>Thr        | gct<br>Ala        | gag<br>Glu        | cgc<br>Arg        | tct<br>Ser<br>620 | ctg<br>Leu        | gaa<br>Glu        | gac<br>Asp        | gag<br>Glu        | 1872 |
| atc<br>Ile<br>625 | aat<br>Asn        | cgc<br>Arg        | acc<br>Thr        | aca<br>Thr        | gct<br>Ala<br>630 | gaa<br>Glu        | gac<br>Asp        | ctg<br>Leu        | ccc<br>Pro        | atc<br>Ile<br>635 | ttt<br>Phe        | gcc<br>Ala        | acc<br>Thr        | agc<br>Ser        | tac<br>Tyr<br>640 | 1920 |
| att<br>Ile        | gtc<br>Val        | ata<br>Ile        | ttc<br>Phe        | ctg<br>Leu<br>645 | tac<br>Tyr        | atc<br>Ile        | tct<br>Ser        | ctg<br>Leu        | gcc<br>Ala<br>650 | ctg<br>Leu        | ggc<br>Gly        | agc<br>Ser        | tat<br>Tyr        | tcc<br>Ser<br>655 | agc<br>Ser        | 1968 |
| tgg<br>Trp        | agc<br>Ser        | cga<br>Arg        | gtg<br>Val<br>660 | atg<br>Met        | gtg<br>Val        | gac<br>Asp        | tcc<br>Ser        | aag<br>Lys<br>665 | gcc<br>Ala        | acg<br>Thr        | ctg<br>Leu        | ggc<br>Gly        | ctc<br>Leu<br>670 | ggc<br>Gly        | GJÀ<br>aaa        | 2016 |
| gtg<br>Val        | gcc<br>Ala        | gtg<br>Val<br>675 | gtc<br>Val        | ctg<br>Leu        | gga<br>Gly        | gca<br>Ala        | gtc<br>Val<br>680 | atg<br>Met        | gct<br>Ala        | gcc<br>Ala        | atg<br>Met        | ggc<br>Gly<br>685 | ttc<br>Phe        | ttc<br>Phe        | tcc<br>Ser        | 2064 |
| tac<br>Tyr        | ttg<br>Leu<br>690 | ggt<br>Gly        | Ile               | Arg               | tcc<br>Ser        | Ser               | Leu               | gtc<br>Val        | atc<br>Ile        | ctg<br>Leu        | caa<br>Gln<br>700 | Val               | gtt<br>Val        | cct<br>Pro        | ttc<br>Phe        | 2112 |
| ctg<br>Leu<br>705 | gtg<br>Val        | ctg<br>Leu        | tcc<br>Ser        | gtg<br>Val        | ggg<br>Gly<br>710 | gct<br>Ala        | gat<br>Asp        | aac<br>Asn        | atc<br>Ile        | ttc<br>Phe<br>715 | atc<br>Ile        | ttt<br>Phe        | gtt<br>Val        | ctc<br>Leu        | gag<br>Glu<br>720 | 2160 |
| tac<br>Tyr        | cag<br>Gln        | agg<br>Arg        | ctg<br>Leu        | ccc<br>Pro<br>725 | cgg<br>Arg        | agg<br>Arg        | cct<br>Pro        | GJA<br>aaa        | gag<br>Glu<br>730 | cca<br>Pro        | cga<br>Arg        | gag<br>Glu        | gtc<br>Val        | cac<br>His<br>735 | att<br>Ile        | 2208 |
| ggg<br>Gly        | cga<br>Arg        | gcc<br>Ala        | cta<br>Leu<br>740 | ggc<br>Gly        | agg<br>Arg        | gtg<br>Val        | gct<br>Ala        | ccc<br>Pro<br>745 | agc<br>Ser        | atg<br>Met        | ctg<br>Leu        | ttg<br>Leu        | tgc<br>Cys<br>750 | agc<br>Ser        | ctc<br>Leu        | 2256 |
| tct<br>Ser        | gag<br>Glu        | gcc<br>Ala<br>755 | atc<br>Ile        | tgc<br>Cys        | ttc<br>Phe        | Phe               | cta<br>Leu<br>760 | Gly<br>aga        | gcc<br>Ala        | ctg<br>Leu        | acc<br>Thr        | ccc<br>Pro<br>765 | atg<br>Met        | cca<br>Pro        | gct<br>Ala        | 2304 |
| gtg<br>Val        | cgg<br>Arg        | acc<br>Thr        | ttt<br>Phe        | gcc<br>Ala        | ctg<br>Leu        | acc<br>Thr        | tct<br>Ser        | ggc<br>Gly        | ctt<br>Leu        | gca<br>Ala        | gtg<br>Val        | atc<br>Ile        | ctt<br>Leu        | gac<br>Asp        | ttc<br>Phe        | 2352 |

770 775 780

| ctc<br>Leu<br>785 | ctg<br>Leu        | cag<br>Gln        | atg<br>Met        | tca<br>Ser        | gcc<br>Ala<br>790 | ttt<br>Phe        | gtg<br>Val        | gcc<br>Ala        | ctg<br>Leu        | ctc<br>Leu<br>795 | tcc<br>Ser        | ctg<br>Leu        | gac<br>Asp        | agc<br>Ser        | aag<br>Lys<br>800 | 2400 |
|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|------|
| agg<br>Arg        | cag<br>Gln        | gag<br>Glu        | gcc<br>Ala        | tcc<br>Ser<br>805 | cgg<br>Arg        | ttg<br>Leu        | gac<br>Asp        | gtc<br>Val        | tgc<br>Cys<br>810 | tgc<br>Cys        | tgt<br>Cys        | gtc<br>Val        | aag<br>Lys        | ccc<br>Pro<br>815 | cag<br>Gln        | 2448 |
| gag<br>Glu        | ctg<br>Leu        | ccc<br>Pro        | ccg<br>Pro<br>820 | cct<br>Pro        | ggc<br>Gly        | cag<br>Gln        | gga<br>Gly        | gag<br>Glu<br>825 | ggg<br>ggg        | ctc<br>Leu        | ctg<br>Leu        | ctt<br>Leu        | ggc<br>Gly<br>830 | ttc<br>Phe        | ttc<br>Phe        | 2496 |
| caa<br>Gln        | aag<br>Lys        | gct<br>Ala<br>835 | tat<br>Tyr        | gcc<br>Ala        | ccc<br>Pro        | ttc<br>Phe        | ctg<br>Leu<br>840 | ctg<br>Leu        | cac<br>His        | tgg<br>Trp        | atc<br>Ile        | act<br>Thr<br>845 | cga<br>Arg        | ggt<br>Gly        | gtt<br>Val        | 2544 |
| gtg<br>Val        | ctg<br>Leu<br>850 | ctg<br>Leu        | ctg<br>Leu        | ttt<br>Phe        | ctc<br>Leu        | gcc<br>Ala<br>855 | ctg<br>Leu        | ttc<br>Phe        | gga<br>Gly        | gtg<br>Val        | agc<br>Ser<br>860 | ctc<br>Leu        | tac<br>Tyr        | tcc<br>Ser        | atg<br>Met        | 2592 |
|                   |                   |                   |                   |                   |                   |                   |                   | cag<br>Gln        |                   |                   |                   |                   |                   |                   |                   | 2640 |
| tcg<br>Ser        | tac<br>Tyr        | ctg<br>Leu        | ctt<br>Leu        | gac<br>Asp<br>885 | tat<br>Tyr        | ttc<br>Phe        | ctc<br>Leu        | ttt<br>Phe        | ctg<br>Leu<br>890 | aac<br>Asn        | cgc<br>Arg        | tac<br>Tyr        | ttc<br>Phe        | gag<br>Glu<br>895 | gtg<br>Val        | 2688 |
| GJÀ<br>aaa        | gcc<br>Ala        | ccg<br>Pro        | gtg<br>Val<br>900 | tac<br>Tyr        | ttt<br>Phe        | gtt<br>Val        | acc<br>Thr        | acc<br>Thr<br>905 | ttg<br>Leu        | ggc               | tac<br>Tyr        | aac<br>Asn        | ttc<br>Phe<br>910 | tcc<br>Ser        | agc<br>Ser        | 2736 |
| gag<br>Glu        | gct<br>Ala        | ggg<br>Gly<br>915 | atg<br>Met        | aat<br>Asn        | gcc<br>Ala        | atc<br>Ile        | tgc<br>Cys<br>920 | tcc<br>Ser        | agt<br>Ser        | gca<br>Ala        | ggc               | tgc<br>Cys<br>925 | aac<br>Asn        | aac<br>Asn        | ttc<br>Phe        | 2784 |
| tcc<br>Ser        | ttc<br>Phe<br>930 | Thr               | cag<br>Gln        | aag<br>Lys        | atc<br>Ile        | cag<br>Gln<br>935 | tat<br>Tyr        | gcc<br>Ala        | aca<br>Thr        | gag<br>Glu        | ttc<br>Phe<br>940 | cct<br>Pro        | gag<br>Glu        | cag<br>Gln        | tct<br>Ser        | 2832 |
| tac<br>Tyr<br>945 | ctg<br>Leu        | gcc<br>Ala        | atc<br>Ile        | cct<br>Pro        | gcc<br>Ala<br>950 | tcc<br>Ser        | tcc<br>Ser        | tgg<br>Trp        | gtg<br>Val        | gat<br>Asp<br>955 | Asp               | ttc<br>Phe        | att<br>Ile        | gac<br>Asp        | tgg<br>Trp<br>960 | 2880 |
|                   |                   |                   |                   |                   |                   |                   |                   | ctt<br>Leu        |                   |                   |                   |                   |                   |                   | Lys               | 2928 |
| gac<br>Asp        | aag<br>Lys        | ttc<br>Phe        | tgc<br>Cys<br>980 | Pro               | tcg<br>Ser        | acc<br>Thr        | gtc<br>Val        | aac<br>Asn<br>985 | Ser               | ctg<br>Leu        | aac<br>Asn        | tgc<br>Cys        | cta<br>Leu<br>990 | Lys               | aac<br>Asn        | 2976 |
|                   |                   |                   | Ile               |                   |                   |                   |                   | gt<br>Va          |                   |                   |                   | r Va              |                   | _                 | ag ttc<br>In Phe  | 3024 |
|                   | aag<br>Lys<br>101 | Ту                |                   |                   | c tg<br>o Tr      | p Ph              |                   | tg a<br>eu A      | _                 |                   | rg F              |                   |                   |                   |                   | 3069 |

| tgt<br>Cys | ccc<br>Pro<br>1025   | aaa<br>Lys | ggc<br>Gly     | ggc<br>Gly | ctg<br>Leu | gca<br>Ala<br>1030 | gca<br>Ala | tac<br>Tyr | agc<br>Ser   | acc<br>Thr | tct<br>Ser<br>1035 | gtg<br>Val | aac<br>Asn | ttg<br>Leu | 3114 |
|------------|----------------------|------------|----------------|------------|------------|--------------------|------------|------------|--------------|------------|--------------------|------------|------------|------------|------|
|            | tca<br>Ser<br>1040   | gat<br>Asp | ggc<br>Gly     | cag<br>Gln | gtt<br>Val | tta<br>Leu<br>1045 | gcc<br>Ala | tcc<br>Ser | agg<br>Arg   | ttc<br>Phe | atg<br>Met<br>1050 | gcc<br>Ala | tat<br>Tyr | cac<br>His | 3159 |
| aag<br>Lys | ccc<br>Pro<br>1055   | ctg<br>Leu | aaa<br>Lys     | aac<br>Asn | tca<br>Ser | cag<br>Gln<br>1060 | gat<br>Asp | tac<br>Tyr | aca<br>Thr   | gaa<br>Glu | gct<br>Ala<br>1065 | ctg<br>Leu | cgg<br>Arg | gca<br>Ala | 3204 |
| _          | cga<br>Arg<br>1070   | gag<br>Glu | ctg<br>Leu     | gca<br>Ala | gcc<br>Ala | aac<br>Asn<br>1075 | atc<br>Ile | act<br>Thr | gct<br>Ala   | gac<br>Asp | ctg<br>Leu<br>1080 | cgg<br>Arg | aaa<br>Lys | gtg<br>Val | 3249 |
|            | gga<br>Gly<br>1085   | Thr        | gac<br>Asp     | ccg<br>Pro | gct<br>Ala | ttt<br>Phe<br>1090 | gag<br>Glu | gtc<br>Val | ttc<br>Phe   | ccc<br>Pro | tac<br>Tyr<br>1095 | acg<br>Thr | atc<br>Ile | acc<br>Thr | 3294 |
|            | gtg<br>Val<br>1100   | Phe        | tat<br>Tyr     | gag<br>Glu | cag<br>Gln | tac<br>Tyr<br>1105 | ctg<br>Leu | acc<br>Thr | atc<br>Ile   | ctc<br>Leu | cct<br>Pro<br>1110 | gag<br>Glu | Gly<br>aaa | ctc<br>Leu | 3339 |
| ttc<br>Phe | atg<br>Met<br>1115   | Leu        | agc<br>Ser     | ctc<br>Leu | tgc<br>Cys | ctt<br>Leu<br>1120 | gtg<br>Val | ccc<br>Pro | acc<br>Thr   | ttc<br>Phe | gct<br>Ala<br>1125 | gtc<br>Val | tcc<br>Ser | tgc<br>Cys | 3384 |
| ctc<br>Leu | ctg<br>Leu<br>1130   | Leu        | ggc            | ctg<br>Leu | gac<br>Asp | ctg<br>Leu<br>1135 | cgc<br>Arg | tcc<br>Ser | ggc          | ctc<br>Leu | ctc<br>Leu<br>1140 | Asn        | ctg<br>Leu | ctc<br>Leu | 3429 |
| tcc<br>Ser | att<br>Ile<br>1145   | Val        | atg<br>Met     | atc<br>Ile | ctc<br>Leu | gtg<br>Val<br>1150 | gac<br>Asp | act<br>Thr | gtc<br>Val   | ggc<br>Gly | ttc<br>Phe<br>1155 | Met        | gcc<br>Ala | ctg<br>Leu | 3474 |
| tgg<br>Trp | gac<br>Asp<br>1160   | Ile        | agt<br>Ser     | tac<br>Tyr | aat<br>Asn | gct<br>Ala<br>1165 | Val        | tcc<br>Ser | ctc<br>Leu   | atc<br>Ile | aac<br>Asn<br>1170 | Leu        | gtc<br>Val | tcg<br>Ser | 3519 |
|            | gtg<br>Val<br>1175   | Gly        | atg<br>Met     | tct<br>Ser | gtg<br>Val | gag<br>Glu<br>1180 | Phe        | gtg<br>Val | tcc<br>Ser   | cac<br>His | att<br>Ile<br>1185 | Thr        | cgc<br>Arg | tcc<br>Ser | 3564 |
|            | gcc<br>Ala<br>1190   | Ile        |                |            |            | ccc<br>Pro<br>1195 | Thr        | tgg<br>Trp | ctg<br>Leu   | gag<br>Glu | agg<br>Arg<br>1200 | Ala        | aaa<br>Lys | gag<br>Glu | 3609 |
|            | acc<br>Thr<br>1205   | Il∈        | tct<br>Ser     | atg<br>Met | gga<br>Gly | agt<br>Ser<br>1210 | Ala        | gtg<br>Val | ttt.<br>Phe  | gca<br>Ala | ggt<br>Gly<br>1215 | Val        | gcc<br>Ala | atg<br>Met | 3654 |
| acc<br>Thi | aac<br>Asn<br>1220   | Leu        | g cct<br>L Pro | ggc<br>Gly | atc<br>Ile | ctt<br>Leu<br>1225 | Val        | cto<br>Lei | ı Gly        | cto<br>Leu | gcc<br>Ala<br>1230 | Lys        | gcc<br>Ala | cag<br>Gln | 3699 |
| cto<br>Lev | att<br>i Ile<br>123! | Glr        | g ato<br>n Ile | ttc<br>Phe | tto<br>Phe | ttc<br>Phe<br>1240 | Arg        | cto<br>Lei | aac<br>1 Asr | cto<br>Lev | ctg<br>Leu<br>1245 | I1e        | act<br>Thr | ctg<br>Leu | 3744 |

| ~ | ggc<br>Gly<br>1250 | _ | _ |   |   | ttg<br>Leu<br>1255 | _ |   | -   |   | _ |       |   | 3789 |
|---|--------------------|---|---|---|---|--------------------|---|---|-----|---|---|-------|---|------|
|   |                    |   |   | - | - | aac<br>Asn<br>1270 | _ | _ | _   | - | - | <br>_ | _ | 3834 |
|   |                    |   | - | - |   | gca<br>Ala<br>1285 |   | - | _   |   |   |       |   | 3879 |
|   |                    |   |   | - | _ | tcc<br>Ser<br>1300 |   | - | -   |   |   | -     |   | 3924 |
|   | agc<br>Ser<br>1310 |   |   |   |   | atc<br>Ile<br>1315 |   |   | _   |   | _ | -     |   | 3969 |
|   | ttg<br>Leu<br>1325 |   |   |   |   | cgg<br>Arg<br>1330 | _ |   | tga |   |   |       |   | 3999 |

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Cys Ala Phe Tyr Asp Glu Cys Gly Lys Asn Pro Glu Leu Ser Gly Ser 35 40 45

Leu Met Thr Leu Ser Asn Val Ser Cys Leu Ser Asn Thr Pro Ala Arg 50 55 60

Lys Ile Thr Gly Asp His Leu Ile Leu Leu Gln Lys Ile Cys Pro Arg 70 75 80

Leu Tyr Thr Gly Pro Asn Thr Gln Ala Cys Cys Ser Ala Lys Gln Leu 85 90 95

Val Ser Leu Glu Ala Ser Leu Ser Ile Thr Lys Ala Leu Leu Thr Arg Cys Pro Ala Cys Ser Asp Asn Phe Val Asn Leu His Cys His Asn Thr Cys Ser Pro Asn Gln Ser Leu Phe Ile Asn Val Thr Arg Val Ala Gln Leu Gly Ala Gly Gln Leu Pro Ala Val Val Ala Tyr Glu Ala Phe Tyr Gln His Ser Phe Ala Glu Gln Ser Tyr Asp Ser Cys Ser Arg Val Arg Val Pro Ala Ala Ala Thr Leu Ala Val Gly Thr Met Cys Gly Val Tyr Gly Ser Ala Leu Cys Asn Ala Gln Arg Trp Leu Asn Phe Gln Gly Asp Thr Gly Asn Gly Leu Ala Pro Leu Asp Ile Thr Phe His Leu Leu Glu Pro Gly Gln Ala Val Gly Ser Gly Ile Gln Pro Leu Asn Glu Gly Val Ala Arg Cys Asn Glu Ser Gln Gly Asp Asp Val Ala Thr Cys Ser Cys Gln Asp Cys Ala Ala Ser Cys Pro Ala Ile Ala Arg Pro Gln Ala Leu Asp Ser Thr Phe Tyr Leu Gly Gln Met Pro Gly Ser Leu Val Leu Ile Ile Ile Leu Cys Ser Val Phe Ala Val Val Thr Ile Leu Leu Val Gly Phe Arg Val Ala Pro Ala Arg Asp Lys Ser Lys Met Val Asp Pro Lys

Leu Gly Gln Phe Phe Gln Gly Trp Gly Thr Trp Val Ala Ser Trp Pro

Lys Gly Thr Ser Leu Ser Asp Lys Leu Ser Phe Ser Thr His Thr Leu

340 345 350

Leu Thr Ile Leu Val Leu Ser Val Ile Pro Val Val Ala Leu Ala Ala 355 360 365

Gly Leu Val Phe Thr Glu Leu Thr Thr Asp Pro Val Glu Leu Trp Ser 370 375 380

Ala Pro Asn Ser Gln Ala Arg Ser Glu Lys Ala Phe His Asp Gln His 385 390 395 400

Phe Gly Pro Phe Phe Arg Thr Asn Gln Val Ile Leu Thr Ala Pro Asn 405 410 415

Arg Ser Ser Tyr Arg Tyr Asp Ser Leu Leu Gly Pro Lys Asn Phe 420 425 430

Ser Gly Ile Leu Asp Leu Asp Leu Leu Leu Glu Leu Glu Leu Gln 435

Glu Arg Leu Arg His Leu Gln Val Trp Ser Pro Glu Ala Gln Arg Asn 450 455 460

Ile Ser Leu Gln AspIle Cys Tyr Ala Pro Leu Asn Pro Asp Asn Thr465470475

Ser Leu Tyr Asp Cys Cys Ile Asn Ser Leu Leu Gln Tyr Phe Gln Asn 485 490 495

Asn Arg Thr Leu Leu Leu Leu Thr Ala Asn Gln Thr Leu Met Gly Gln 500 505 510

Thr Ser Gln Val Asp Trp Lys Asp His Phe Leu Tyr Cys Ala Asn Ala 515 520 525

Pro Leu Thr Phe Lys Asp Gly Thr Ala Leu Ala Leu Ser Cys Met Ala 530 540

Asp Tyr Gly Ala Pro Val Phe Pro Phe Leu Ala Ile Gly Gly Tyr Lys 545 550 555 560

Gly Lys Asp Tyr Ser Glu Ala Glu Ala Leu Ile Met Thr Phe Ser Leu 565 570 575

Asn Asn Tyr Pro Ala Gly Asp Pro Arg Leu Ala Gln Ala Lys Leu Trp 580 585 590

Glu Glu Ala Phe Leu Glu Glu Met Arg Ala Phe Gln Arg Arg Met Ala 595 600 605

Gly Met Phe Gln Val Thr Phe Thr Ala Glu Arg Ser Leu Glu Asp Glu 610 620

Ile Asn Arg Thr Thr Ala Glu Asp Leu Pro Ile Phe Ala Thr Ser Tyr 625 630 635 640

Ile Val Ile Phe Leu Tyr Ile Ser Leu Ala Leu Gly Ser Tyr Ser Ser 645 650 655

Trp Ser Arg Val Met Val Asp Ser Lys Ala Thr Leu Gly Leu Gly Gly 660 665 670

Val Ala Val Val Leu Gly Ala Val Met Ala Ala Met Gly Phe Phe Ser 675 680 685

Tyr Leu Gly Ile Arg Ser Ser Leu Val Ile Leu Gln Val Val Pro Phe 690 695 700

Leu Val Leu Ser Val Gly Ala Asp Asn Ile Phe Ile Phe Val Leu Glu 705 710 715 720

Tyr Gln Arg Leu Pro Arg Arg Pro Gly Glu Pro Arg Glu Val His Ile 725 730 735

Gly Arg Ala Leu Gly Arg Val Ala Pro Ser Met Leu Leu Cys Ser Leu 740 745 750

Ser Glu Ala Ile Cys Phe Phe Leu Gly Ala Leu Thr Pro Met Pro Ala 755 760 765

Val Arg Thr Phe Ala Leu Thr Ser Gly Leu Ala Val Ile Leu Asp Phe 770 780

Leu Leu Gln Met Ser Ala Phe Val Ala Leu Leu Ser Leu Asp Ser Lys 785 790 795 800

Arg Gln Glu Ala Ser Arg Leu Asp Val Cys Cys Cys Val Lys Pro Gln 805 810 815

Glu Leu Pro Pro Pro Gly Gln Gly Glu Gly Leu Leu Gly Phe Phe 820 825 830

- Gln Lys Ala Tyr Ala Pro Phe Leu Leu His Trp Ile Thr Arg Gly Val 835 840 845
- Val Leu Leu Phe Leu Ala Leu Phe Gly Val Ser Leu Tyr Ser Met 850 855
- Cys His Ile Ser Val Gly Leu Asp Gln Glu Leu Ala Leu Pro Lys Asp 865 870 875 880
- Ser Tyr Leu Leu Asp Tyr Phe Leu Phe Leu Asn Arg Tyr Phe Glu Val 885 890 895
- Gly Ala Pro Val Tyr Phe Val Thr Thr Leu Gly Tyr Asn Phe Ser Ser 900 905
- Glu Ala Gly Met Asn Ala Ile Cys Ser Ser Ala Gly Cys Asn Asn Phe 915 920 925
- Ser Phe Thr Gln Lys Ile Gln Tyr Ala Thr Glu Phe Pro Glu Gln Ser 930 935 940
- Tyr Leu Ala Ile Pro Ala Ser Ser Trp Val Asp Asp Phe Ile Asp Trp 945 950 955 960
- Leu Thr Pro Ser Ser Cys Cys Arg Leu Tyr Ile Ser Gly Pro Asn Lys 965 970 975
- Asp Lys Phe Cys Pro Ser Thr Val Asn Ser Leu Asn Cys Leu Lys Asn 980 985
- Cys Met Ser Ile Thr Met Gly Ser Val Arg Pro Ser Val Glu Gln Phe 995 1000 1005
- His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Arg Pro Asn Ile Lys 1010 1015
- Cys Pro Lys Gly Gly Leu Ala Ala Tyr Ser Thr Ser Val Asn Leu 1025 1035
- Thr Ser Asp Gly Gln Val Leu Ala Ser Arg Phe Met Ala Tyr His 1040 1045 1050
- Lys Pro Leu Lys Asn Ser Gln Asp Tyr Thr Glu Ala Leu Arg Ala 1055 1060 1065

- Ala Arg Glu Leu Ala Ala Asn Ile Thr Ala Asp Leu Arg Lys Val Pro Gly Thr Asp Pro Ala Phe Glu Val Phe Pro Tyr Thr Ile Thr Asn Val Phe Tyr Glu Gln Tyr Leu Thr Ile Leu Pro Glu Gly Leu Phe Met Leu Ser Leu Cys Leu Val Pro Thr Phe Ala Val Ser Cys Leu Leu Leu Gly Leu Asp Leu Arg Ser Gly Leu Leu Asn Leu Leu Ser Ile Val Met Ile Leu Val Asp Thr Val Gly Phe Met Ala Leu Trp Asp Ile Ser Tyr Asn Ala Val Ser Leu Ile Asn Leu Val Ser Ala Val Gly Met Ser Val Glu Phe Val Ser His Ile Thr Arg Ser Phe Ala Ile Ser Thr Lys Pro Thr Trp Leu Glu Arg Ala Lys Glu Ala Thr Ile Ser Met Gly Ser Ala Val Phe Ala Gly Val Ala Met Thr Asn Leu Pro Gly Ile Leu Val Leu Gly Leu Ala Lys Ala Gln Leu Ile Gln Ile Phe Phe Phe Arg Leu Asn Leu Leu Ile Thr Leu
  - Leu Gly Leu Leu His Gly Leu Val Phe Leu Pro Val Ile Leu Ser 1250 1255 1260
  - Tyr Val Gly Pro Asp Val Asn Pro Ala Leu Ala Leu Glu Gln Lys 1265 1270 1275
  - Arg Ala Glu Glu Ala Val Ala Ala Val Met Val Ala Ser Cys Pro 1280 1285 1290
  - Asn His Pro Ser Arg Val Ser Thr Ala Asp Asn Ile Tyr Val Asn

1295 1300 1305

His Ser Phe Glu Gly Ser Ile Lys Gly Ala Gly Ala Ile Ser Asn 1310 1315 1320

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Arg Asp Pro Gly Gln Leu Pro Ala Val Val Ala Tyr Glu Ala Phe Tyr

| 145            | 150 |   | 155                                       | 160 |
|----------------|-----|---|---|-----|
| ~ ~            |     |   | tcc tgt agc cgg<br>Ser Cys Ser Arg        |     |
| Ile Pro Ala Al |     |   | agc atg tgt gga<br>Ser Met Cys Gly<br>190 |     |
|                | -   |   | ctc aac ttc caa<br>Leu Asn Phe Gln<br>205 |     |
|                |     |   | acc ttc cac ctc<br>Thr Phe His Leu<br>220 |     |
|                |     |   | cca ctg gat ggg<br>Pro Leu Asp Gly<br>235 |     |
| _              |     |   | tcg gca gcc tgt<br>Ser Ala Ala Cys        | -   |
| Gln Asp Cys A  | -   | _ | cct ccg ccc ccg<br>Pro Pro Pro Pro<br>270 |     |
| -              |     |   | ggc tgg ctg gct<br>Gly Trp Leu Ala<br>285 |     |
|                |     | _ | tct gtt gtc ctt<br>Ser Val Val Leu<br>300 |     |
|                |     | _ | aag aca gca ggc<br>Lys Thr Ala Gly<br>315 |     |
|                | -   |   | ttc tca cct cac<br>Phe Ser Pro His        |     |
| Leu Gly Arg Pl |     |   | agg gtg gcc tca<br>Arg Val Ala Ser<br>350 |     |
|                |     |   | gtg ata gcc ttg<br>Val Ile Ala Leu<br>365 |     |
|                | _   |   | cct gtg gaa ctg<br>Pro Val Glu Leu<br>380 |     |
|                |     |   | gct ttc cat gac<br>Ala Phe His Asp<br>395 |     |

| ttt<br>Phe        | ggc<br>Gly        | ccc<br>Pro        | ttc<br>Phe        | ttc<br>Phe<br>405 | cga<br>Arg        | acc<br>Thr        | aac<br>Asn        | cag<br>Gln        | att<br>Ile<br>410 | ttt<br>Phe        | gtg<br>Val        | aca<br>Thr        | gct<br>Ala        | aag<br>Lys<br>415 | aac<br>Asn        | 1248 |
|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|------|
| agg<br>Arg        | tcc<br>Ser        | agc<br>Ser        | tac<br>Tyr<br>420 | aag<br>Lys        | tac<br>Tyr        | gac<br>Asp        | tcc<br>Ser        | ctg<br>Leu<br>425 | ctg<br>Leu        | cta<br>Leu        | Gly<br>ggg        | ccc<br>Pro        | aag<br>Lys<br>430 | aac<br>Asn        | ttc<br>Phe        | 1296 |
| agt<br>Ser        | Gly<br>ggg        | atc<br>Ile<br>435 | cta<br>Leu        | tcc<br>Ser        | ctg<br>Leu        | gac<br>Asp        | ttg<br>Leu<br>440 | ctg<br>Leu        | cag<br>Gln        | gag<br>Glu        | ctg<br>Leu        | ttg<br>Leu<br>445 | gag<br>Glu        | cta<br>Leu        | cag<br>Gln        | 1344 |
| gag<br>Glu        | aga<br>Arg<br>450 | ctt<br>Leu        | cga<br>Arg        | cac<br>His        | ctg<br>Leu        | caa<br>Gln<br>455 | gtg<br>Val        | tgg<br>Trp        | tcc<br>Ser        | cat<br>His        | gag<br>Glu<br>460 | gca<br>Ala        | cag<br>Gln        | cgc<br>Arg        | aac<br>Asn        | 1392 |
| atc<br>Ile<br>465 | tcc<br>Ser        | ctc<br>Leu        | cag<br>Gln        | gac<br>Asp        | atc<br>Ile<br>470 | tgc<br>Cys        | tat<br>Tyr        | gct<br>Ala        | ccc<br>Pro        | ctc<br>Leu<br>475 | aac<br>Asn        | ccg<br>Pro        | cat<br>His        | aac<br>Asn        | acc<br>Thr<br>480 | 1440 |
| agc<br>Ser        | ctc<br>Leu        | act<br>Thr        | gac<br>Asp        | tgc<br>Cys<br>485 | tgt<br>Cys        | gtc<br>Val        | aac<br>Asn        | agc<br>Ser        | ctc<br>Leu<br>490 | ctt<br>Leu        | caa<br>Gln        | tac<br>Tyr        | ttc<br>Phe        | cag<br>Gln<br>495 | aac<br>Asn        | 1488 |
| aac<br>Asn        | cac<br>His        | aca<br>Thr        | ctc<br>Leu<br>500 | ctg<br>Leu        | ctg<br>Leu        | ctc<br>Leu        | aca<br>Thr        | gcc<br>Ala<br>505 | aat<br>Asn        | cag<br>Gln        | act<br>Thr        | ctg<br>Leu        | aat<br>Asn<br>510 | ggc               | cag<br>Gln        | 1536 |
| acc<br>Thr        | tcc<br>Ser        | ctg<br>Leu<br>515 | gtg<br>Val        | gac<br>Asp        | tgg<br>Trp        | aag<br>Lys        | gac<br>Asp<br>520 | cat<br>His        | ttc<br>Phe        | ctc<br>Leu        | tac<br>Tyr        | tgt<br>Cys<br>525 | gcc<br>Ala        | aat<br>Asn        | gcc<br>Ala        | 1584 |
| cct<br>Pro        | ctc<br>Leu<br>530 | acg<br>Thr        | tac<br>Tyr        | aaa<br>Lys        | gat<br>Asp        | ggc<br>Gly<br>535 | aca<br>Thr        | gcc<br>Ala        | ctg<br>Leu        | gcc<br>Ala        | ctg<br>Leu<br>540 | agc<br>Ser        | tgc<br>Cys        | ata<br>Ile        | gct<br>Ala        | 1632 |
| gac<br>Asp<br>545 | Tyr               | ggg               | gca<br>Ala        | cct<br>Pro        | gtc<br>Val<br>550 | ttc<br>Phe        | ccc<br>Pro        | ttc<br>Phe        | ctt<br>Leu        | gct<br>Ala<br>555 | gtt<br>Val        | ggg               | ggc               | tac<br>Tyr        | caa<br>Gln<br>560 | 1680 |
| Gly<br>ggg        | acg<br>Thr        | gac<br>Asp        | tac<br>Tyr        | tcg<br>Ser<br>565 | Glu               | gca<br>Ala        | gaa<br>Glu        | gcc<br>Ala        | ctg<br>Leu<br>570 | Ile               | ata<br>Ile        | acc<br>Thr        | ttc<br>Phe        | tct<br>Ser<br>575 | Ile               | 1728 |
| aat<br>Asn        | aac<br>Asn        | tac<br>Tyr        | ccc<br>Pro<br>580 | Ala               | gat<br>Asp        | gat<br>Asp        | ccc<br>Pro        | cgc<br>Arg<br>585 | Met               | gcc<br>Ala        | cac<br>His        | gcc<br>Ala        | aag<br>Lys<br>590 | Leu               | tgg<br>Trp        | 1776 |
| gag<br>Glu        | gag<br>Glu        | gct<br>Ala<br>595 | Phe               | ttg<br>Lev        | aag<br>Lys        | gaa<br>Glu        | atg<br>Met<br>600 | Gln               | tcc<br>Ser        | tto<br>Phe        | cag<br>Gln        | aga<br>Arg<br>605 | Ser               | aca<br>Thr        | gct<br>Ala        | 1824 |
| gac<br>Asp        | aag<br>Lys<br>610 | Phe               | cag<br>Glr        | att<br>11e        | gcg<br>Ala        | tto<br>Phe<br>615 | e Ser             | gct<br>Ala        | gag<br>Glu        | cgt<br>Arg        | tct<br>Ser<br>620 | Leu               | gag<br>Glu        | gac<br>Asp        | gag<br>Glu        | 1872 |
| atc<br>Ile<br>625 | e Asr             | cgo<br>Arg        | act<br>Thr        | aco<br>Thr        | ato<br>11e<br>630 | e Glr             | gad<br>Asp        | ctg<br>Lev        | g cct<br>L Pro    | gtc<br>Val<br>635 | . Phe             | gcc<br>Ala        | ato<br>lle        | ago<br>Ser        | tac<br>Tyr<br>640 | 1920 |

| ctt<br>Leu        | atc<br>Ile          | gtc<br>Val        | ttc<br>Phe        | ctg<br>Leu<br>645 | tac<br>Tyr        | atc<br>Ile        | tcc<br>Ser        | ctg<br>Leu        | gcc<br>Ala<br>650 | ctg<br>Leu        | ggc<br>Gly        | agc<br>Ser        | tac<br>Tyr        | tcc<br>Ser<br>655 | aga<br>Arg        | 1968 |
|-------------------|---------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|------|
| tgg<br>Trp        | agc<br>Ser          | cga<br>Arg        | gtt<br>Val<br>660 | gcg<br>Ala        | gtg<br>Val        | gat<br>Asp        | tcc<br>Ser        | aag<br>Lys<br>665 | gct<br>Ala        | act<br>Thr        | ctg<br>Leu        | ggc<br>Gly        | cta<br>Leu<br>670 | ggt<br>Gly        | Gly<br>ggg        | 2016 |
| gtg<br>Val        | gct<br>Ala          | gtt<br>Val<br>675 | gtg<br>Val        | ctg<br>Leu        | gga<br>Gly        | gca<br>Ala        | gtc<br>Val<br>680 | gtc<br>Val        | gct<br>Ala        | gcc<br>Ala        | atg<br>Met        | ggc<br>Gly<br>685 | ttc<br>Phe        | tac<br>Tyr        | tcc<br>Ser        | 2064 |
| tac<br>Tyr        | ctg<br>Leu<br>690   | ggt<br>Gly        | gtc<br>Val        | ccc<br>Pro        | tcc<br>Ser        | tct<br>Ser<br>695 | ctg<br>Leu        | gtc<br>Val        | atc<br>Ile        | att<br>Ile        | caa<br>Gln<br>700 | gtg<br>Val        | gta<br>Val        | cct<br>Pro        | ttc<br>Phe        | 2112 |
| ctg<br>Leu<br>705 | gtg<br>Val          | ctg<br>Leu        | gct<br>Ala        | gtg<br>Val        | gga<br>Gly<br>710 | gct<br>Ala        | gac<br>Asp        | aac<br>Asn        | atc<br>Ile        | ttc<br>Phe<br>715 | atc<br>Ile        | ttt<br>Phe        | gtt<br>Val        | ctt<br>Leu        | gag<br>Glu<br>720 | 2160 |
| tac<br>Tyr        | cag<br>Gln          | agg<br>Arg        | ctg<br>Leu        | cct<br>Pro<br>725 | agg<br>Arg        | atg<br>Met        | ccc<br>Pro        | ggg               | gag<br>Glu<br>730 | cag<br>Gln        | cga<br>Arg        | gag<br>Glu        | gct<br>Ala        | cac<br>His<br>735 | att<br>Ile        | 2208 |
| ggc<br>Gly        | cgc<br>Arg          | acc<br>Thr        | ctg<br>Leu<br>740 | ggt<br>Gly        | agt<br>Ser        | gtg<br>Val        | gcc<br>Ala        | ccc<br>Pro<br>745 | agc<br>Ser        | atg<br>Met        | ctg<br>Leu        | ctg<br>Leu        | tgc<br>Cys<br>750 | agc<br>Ser        | ctc<br>Leu        | 2256 |
| tct<br>Ser        | gag<br>Glu          | gcc<br>Ala<br>755 | atc<br>Ile        | tgc<br>Cys        | ttc<br>Phe        | ttt<br>Phe        | cta<br>Leu<br>760 | Gly               | gcc<br>Ala        | ctg<br>Leu        | acc<br>Thr        | tcc<br>Ser<br>765 | atg<br>Met        | cca<br>Pro        | gct<br>Ala        | 2304 |
| gtg<br>Val        | agg<br>Arg<br>770   | Thr               | ttt<br>Phe        | gcc<br>Ala        | ttg<br>Leu        | acc<br>Thr<br>775 | tct<br>Ser        | ggc               | tta<br>Leu        | gca<br>Ala        | atc<br>Ile<br>780 | atc<br>Ile        | ttt<br>Phe        | gac<br>Asp        | ttc<br>Phe        | 2352 |
| ctg<br>Leu<br>785 | Leu                 | cag<br>Gln        | atg<br>Met        | aca<br>Thr        | gcc<br>Ala<br>790 | ttt<br>Phe        | gtg<br>Val        | gcc<br>Ala        | ctg<br>Leu        | ctc<br>Leu<br>795 | tcc<br>Ser        | ctg<br>Leu        | gat<br>Asp        | agc<br>Ser        | aag<br>Lys<br>800 | 2400 |
| agg<br>Arg        | cag<br>Gln          | gag<br>Glu        | gcc<br>Ala        | tct<br>Ser<br>805 | Arg               | ccc<br>Pro        | gac<br>Asp        | gtc<br>Val        | gtg<br>Val<br>810 | tgc<br>Cys        | tgc<br>Cys        | ttt<br>Phe        | tca<br>Ser        | agc<br>Ser<br>815 | Arg               | 2448 |
| aat<br>Asr        | ctg<br>Leu          | ccc<br>Pro        | cca<br>Pro<br>820 | Pro               | aaa<br>Lys        | caa<br>Gln        | aaa<br>Lys        | gaa<br>Glu<br>825 |                   | ctc<br>Leu        | tta<br>Leu        | ctt<br>Leu        | tgc<br>Cys<br>830 | Phe               | ttc<br>Phe        | 2496 |
| cgc<br>Arg        | : aag<br>  Lys      | ata<br>Ile<br>835 | Tyr               | act<br>Thr        | ccc<br>Pro        | ttc<br>Phe        | ctg<br>Leu<br>840 | Leu               | cac<br>His        | aga<br>Arg        | ttc<br>Phe        | atc<br>Ile<br>845 | Arg               | cct<br>Pro        | gtt<br>Val        | 2544 |
| gto<br>Val        | g ctg<br>Leu<br>850 | ı Lev             | cto<br>Leu        | tttı<br>Phe       | ctg<br>Leu        | gtc<br>Val<br>855 | Leu               | ttt<br>Phe        | gga<br>Gly        | gca<br>Ala        | aac<br>Asn<br>860 | Leu               | tac<br>Tyr        | tta<br>Leu        | atg<br>Met        | 2592 |
| tg:<br>Cys<br>865 | s Asr               | ato<br>11e        | ago<br>Ser        | gtg<br>Val        | ggg<br>Gly<br>870 | Leu               | gac<br>Asp        | caç<br>Glr        | g gat<br>n Asp    | ctg<br>Leu<br>875 | Ala               | ctg<br>Leu        | ccc<br>Pro        | aag<br>Lys        | gat<br>Asp<br>880 | 2640 |
| tco               | c tac               | c ctg             | g ata             | a gad             | tac               | : ttc             | cto               | ttt               | ctg               | aac               | cgg               | tac               | : ttg             | gaa               | gtg               | 2688 |

| Ser Tyr Leu Ile Asp Tyr Phe Leu Phe Leu Asn Arg Tyr Leu Glu Val<br>885 890 895  |                              |
|---|------------------------------|
| ggg cct cca gtg tac ttt gac acc acc tca ggc tac aac ttt tcc acc Gly Pro Pro Val Tyr Phe Asp Thr Thr Ser Gly Tyr Asn Phe Ser Thr 900 905 910   | 2736                         |
| gag gca ggc atg aac gcc att tgc tct agt gca ggc tgt gag agc ttc<br>Glu Ala Gly Met Asn Ala Ile Cys Ser Ser Ala Gly Cys Glu Ser Phe<br>915 920 925   | 2784                         |
| tcc cta acc cag aaa atc cag tat gcc agt gaa ttc cct aat cag tct<br>Ser Leu Thr Gln Lys Ile Gln Tyr Ala Ser Glu Phe Pro Asn Gln Ser<br>930 935 940   | 2832                         |
| tat gtg gct att gct gca tcc tcc tgg gta gat gac ttc atc gac tgg Tyr Val Ala Ile Ala Ala Ser Ser Trp Val Asp Asp Phe Ile Asp Trp 945 950 955 960   | 2880                         |
| ctg acc cca tcc tcc tcc tgc tgc cgc att tat acc cgt ggc ccc cat<br>Leu Thr Pro Ser Ser Ser Cys Cys Arg Ile Tyr Thr Arg Gly Pro His<br>965 970 975   | 2928                         |
| aaa gat gag ttc tgt ccc tca acg gat act tcc ttc aac tgt ctc aaa<br>Lys Asp Glu Phe Cys Pro Ser Thr Asp Thr Ser Phe Asn Cys Leu Lys<br>980 985 990   | 2976                         |
| aac tgc atg aac cgc act ctg ggt ccc gtg aga ccc aca aca gaa cag<br>Asn Cys Met Asn Arg Thr Leu Gly Pro Val Arg Pro Thr Thr Glu Glr<br>995 1000 1005   |                              |
|   |                              |
| ttt cat aag tac ctg ccc tgg ttc ctg aat gat acg ccc aac atc Phe His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Thr Pro Asn Ile 1010 1020   | 3069                         |
| Phe His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Thr Pro Asn Ile   | 3069<br>3114                 |
| Phe His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Thr Pro Asn Ile 1010 1015 1020  aga tgt cct aaa ggg ggc cta gca gcg tat aga acc tct gtg aat Arg Cys Pro Lys Gly Gly Leu Ala Ala Tyr Arg Thr Ser Val Asn   |                              |
| Phe His Lys Tyr Leu Pro Trp 1015 Phe Leu Asn Asp Thr 1020 Pro Asn Ile 1010 1020  aga tgt cct aaa ggg ggc cta gca gcg tat aga acc tct gtg aat Arg Cys Pro Lys Gly Gly Leu Ala Ala Tyr Arg Thr 1035 Ser Val Asn 1025 1030 1035  ttg agc tca gat ggc cag att ata gcc tcc cag ttc atg gcc tac Leu Ser Ser Asp Gly Gln Ile Ile Ala Ser Gln Phe Met Ala Tyr   | 3114                         |
| Phe His Lys Tyr Leu Pro Trp 1015  aga tgt cct aaa ggg ggc cta gca gcg tat aga acc tct gtg aat Arg Cys Pro Lys Gly Gly Leu Ala Ala Tyr Arg Thr 1035  ttg agc tca gat ggc cag att at agc tcc cag ttc atg gcc tac Leu Ser Asp Gly Gln Ile 1045  cac aag ccc tta cgg aac tca cag gac ttt aca gaa gct ctc cgg His Lys Pro Leu Arg Asn Ser Gln Asp Phe Thr Glu Ala Leu Arg  | 3114<br>3159                 |
| Phe His Lys Tyr Leu Pro Trp 1015  aga tgt cct aaa ggg ggc cta gca gcg tat aga acc tct gtg aat Arg Cys Pro Lys Gly Gly Leu Ala Ala Tyr Arg Thr 1035  ttg agc tca gat ggc cag att at agc tcc cag ttc atg gcc tac Leu Ser Asp Gly Gln Ile 1045  cac aag ccc tta cgg aac tca cag gac ttt aca gaa gct ctc cgg His Lys Pro Leu Arg Asn Ser Gln Asp Phe Thr Glu Ala Leu Arg 1055  gca tcc cgg ttg cta gca gcc aac atc aca gct gaa cta cgg aag Ala Ser Arg Leu Leu Ala Ala Ala Ala Thr Ala Glu Leu Arg Lys  | 3114<br>3159<br>3204         |
| Phe His Lys Tyr Leu Pro Trp 1015  aga tgt cct aaa ggg ggc cta Arg Cys Pro Lys Gly Gly Leu 1030  ttg agc tca gat ggc cag att 1030  ttg agc tca gat ggc cag att 1030  ttg agc tca gat ggc cag att 1040  cac aag ccc tta cgg aac tca 1040  Fro Asn Ile 1020  repro Asn Ile 1020  fro Asn Ile | 3114<br>3159<br>3204<br>3249 |

|   | 1115               |     |   |   |   | 1120               |   |   |   |     | 1125               |   |   |   |      |
|---|--------------------|-----|---|---|---|--------------------|---|---|---|-----|--------------------|---|---|---|------|
|   |                    |     |   |   |   | gac<br>Asp<br>1135 |   |   |   |     |                    |   |   |   | 3429 |
|   |                    |     |   |   |   | ctc<br>Leu<br>1150 |   |   |   |     |                    |   |   |   | 3474 |
|   |                    | -   |   | _ |   | aat<br>Asn<br>1165 |   | _ |   |     |                    |   |   |   | 3519 |
| _ |                    |     |   | _ |   | gtg<br>Val<br>1180 | - |   |   |     |                    |   |   |   | 3564 |
|   | ttt<br>Phe<br>1190 |     |   |   |   | aag<br>Lys<br>1195 |   |   |   |     |                    |   |   |   | 3609 |
| _ | gct<br>Ala<br>1205 |     |   |   | _ | ggc<br>Gly<br>1210 | - | _ |   |     |                    | _ |   |   | 3654 |
| ~ | acc<br>Thr<br>1220 |     |   |   |   | atc<br>Ile<br>1225 |   |   |   |     |                    |   |   |   | 3699 |
| _ | ctt<br>Leu<br>1235 |     | _ |   |   | ttc<br>Phe<br>1240 |   | _ |   |     |                    | _ |   |   | 3744 |
| _ | _                  |     | _ |   |   | ggc<br>Gly<br>1255 | _ |   |   | _   |                    | - | _ |   | 3789 |
|   | tat<br>Tyr<br>1265 | _   |   |   | _ | gtt<br>Val<br>1270 |   |   | _ | _   | -                  | _ |   |   | 3834 |
|   |                    | _   |   |   | _ | gcc<br>Ala<br>1285 | _ | - |   |     |                    |   | _ |   | 3879 |
| _ |                    |     |   | _ | - | gat<br>Asp<br>1300 | _ |   |   | _   | -                  |   |   |   | 3924 |
|   |                    | Phe |   |   | - | ttt<br>Phe<br>1315 |   |   | _ |     | aat<br>Asn<br>1320 | - | - | _ | 3969 |
| • | tct<br>Ser<br>1325 | _   |   |   | _ | gac<br>Asp<br>1330 |   | _ |   | taa |                    |   |   |   | 4002 |

<210> 12

<211> 1333

<212> PRT

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<400> 12

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Ser Ala Gln Gly Glu Leu Tyr Thr Pro Thr His Lys Ala Gly Phe Cys 20 25 30

Thr Phe Tyr Glu Glu Cys Gly Lys Asn Pro Glu Leu Ser Gly Gly Leu 35 40 45

Thr Ser Leu Ser Asn Ile Ser Cys Leu Ser Asn Thr Pro Ala Arg His 50 55 60

Val Thr Gly Asp His Leu Ala Leu Leu Gln Arg Val Cys Pro Arg Leu 65 70 75 80

Tyr Asn Gly Pro Asn Asp Thr Tyr Ala Cys Cys Ser Thr Lys Gln Leu 85 90 95

Val Ser Leu Asp Ser Ser Leu Ser Ile Thr Lys Ala Leu Leu Thr Arg 100 105 110

Cys Pro Ala Cys Ser Glu Asn Phe Val Ser Ile His Cys His Asn Thr 115 120 125

Cys Ser Pro Asp Gln Ser Leu Phe Ile Asn Val Thr Arg Val Val Gln 130 135 140

Arg Asp Pro Gly Gln Leu Pro Ala Val Val Ala Tyr Glu Ala Phe Tyr 145 150 155 160

Gln Arg Ser Phe Ala Glu Lys Ala Tyr Glu Ser Cys Ser Arg Val Arg 165 170 175

Ile Pro Ala Ala Ala Ser Leu Ala Val Gly Ser Met Cys Gly Val Tyr 180 185 190

Gly Ser Ala Leu Cys Asn Ala Gln Arg Trp Leu Asn Phe Gln Gly Asp 195 200 205 Thr Gly Asn Gly Leu Ala Pro Leu Asp Ile Thr Phe His Leu Leu Glu 220 Pro Gly Gln Ala Leu Ala Asp Gly Met Lys Pro Leu Asp Gly Lys Ile Thr Pro Cys Asn Glu Ser Gln Gly Glu Asp Ser Ala Ala Cys Ser Cys Gln Asp Cys Ala Ala Ser Cys Pro Val Ile Pro Pro Pro Pro Ala Leu 265 Arg Pro Ser Phe Tyr Met Gly Arg Met Pro Gly Trp Leu Ala Leu Ile Ile Ile Phe Thr Ala Val Phe Val Leu Ser Val Val Leu Val Tyr 295 Leu Arg Val Ala Ser Asn Arg Asn Lys Asn Lys Thr Ala Gly Ser Gln 310 315 Glu Ala Pro Asn Leu Pro Arg Lys Arg Phe Ser Pro His Thr Val 330 Leu Gly Arg Phe Phe Glu Ser Trp Gly Thr Arg Val Ala Ser Trp Pro 340 345 Leu Thr Val Leu Ala Leu Ser Phe Ile Val Val Ile Ala Leu Ser Val 355 360 365 Gly Leu Thr Phe Ile Glu Leu Thr Thr Asp Pro Val Glu Leu Trp Ser 370 375 Ala Pro Lys Ser Gln Ala Arg Lys Glu Lys Ala Phe His Asp Glu His 390 395 Phe Gly Pro Phe Phe Arg Thr Asn Gln Ile Phe Val Thr Ala Lys Asn 410 405 Arg Ser Ser Tyr Lys Tyr Asp Ser Leu Leu Leu Gly Pro Lys Asn Phe 425 420 Ser Gly Ile Leu Ser Leu Asp Leu Leu Gln Glu Leu Leu Gln Leu Gln 435 440

Glu Arg Leu Arg His Leu Gln Val Trp Ser His Glu Ala Gln Arg Asn Ile Ser Leu Gln Asp Ile Cys Tyr Ala Pro Leu Asn Pro His Asn Thr Ser Leu Thr Asp Cys Cys Val Asn Ser Leu Leu Gln Tyr Phe Gln Asn Asn His Thr Leu Leu Leu Thr Ala Asn Gln Thr Leu Asn Gly Gln Thr Ser Leu Val Asp Trp Lys Asp His Phe Leu Tyr Cys Ala Asn Ala Pro Leu Thr Tyr Lys Asp Gly Thr Ala Leu Ala Leu Ser Cys Ile Ala Asp Tyr Gly Ala Pro Val Phe Pro Phe Leu Ala Val Gly Gly Tyr Gln Gly Thr Asp Tyr Ser Glu Ala Glu Ala Leu Ile Ile Thr Phe Ser Ile Asn Asn Tyr Pro Ala Asp Asp Pro Arg Met Ala His Ala Lys Leu Trp Glu Glu Ala Phe Leu Lys Glu Met Gln Ser Phe Gln Arg Ser Thr Ala Asp Lys Phe Gln Ile Ala Phe Ser Ala Glu Arg Ser Leu Glu Asp Glu Ile Asn Arg Thr Thr Ile Gln Asp Leu Pro Val Phe Ala Ile Ser Tyr Leu Ile Val Phe Leu Tyr Ile Ser Leu Ala Leu Gly Ser Tyr Ser Arg Trp Ser Arg Val Ala Val Asp Ser Lys Ala Thr Leu Gly Leu Gly Gly Val Ala Val Val Leu Gly Ala Val Val Ala Ala Met Gly Phe Tyr Ser 

Tyr Leu Gly Val Pro Ser Ser Leu Val Ile Ile Gln Val Val Pro Phe

690 695 700

| Leu<br>705      | Val        | Leu          | Ala        | Val          | Gly<br>710 | Ala        | Asp          | Asn        | Ile          | Phe<br>715 | Ile        | Phe          | Val        | Leu        | Glu<br>720 |
|-----------------|------------|--------------|------------|--------------|------------|------------|--------------|------------|--------------|------------|------------|--------------|------------|------------|------------|
| Tyr             | Gln        | Arg          | Leu        | Pro<br>725   | Arg        | Met        | Pro          | Gly        | Glu<br>730   | Gln        | Arg        | Glu          | Ala        | His<br>735 | Ile        |
| Gly             | Arg        | Thr          | Leu<br>740 | Gly          | Ser        | Val        | Ala          | Pro<br>745 | Ser          | Met        | Leu        | Leu          | Cys<br>750 | Ser        | Leu        |
| Ser             | Glu        | Ala<br>755   | Ile        | Cys          | Phe        | Phe        | Leu<br>760   | Gly        | Ala          | Leu        | Thr        | Ser<br>765   | Met        | Pro        | Ala        |
| Val             | Arg<br>770 | Thr          | Phe        | Ala          | Leu        | Thr<br>775 | Ser          | Gly        | Leu          | Ala        | Ile<br>780 | Ile          | Phe        | Asp        | Phe        |
| Leu<br>785      | Leu        | Gln          | Met        | Thr          | Ala<br>790 | Phe        | Val          | Ala        | Leu          | Leu<br>795 | Ser        | Leu          | Asp        | Ser        | Lys<br>800 |
| Arg             | Gln        | Glu          | Ala        | Ser<br>805   | Arg        | Pro        | Asp          | Val        | Val<br>810   |            | Cys        | Phe          | Ser        | Ser<br>815 | Arg        |
| Asn             | Leu        | Pro          | Pro<br>820 | Pro          | Lys        | Gln        | Lys          | Glu<br>825 |              | Leu        | Leu        | Leu          | Cys<br>830 | Phe        | Phe        |
| Arg             | Lys        | Ile<br>835   | Tyr        | Thr          | Pro        | Phe        | Leu<br>840   |            | His          | Arg        | Phe        | Ile<br>845   |            | Pro        | Val        |
| Val             | Leu<br>850 |              | . Leu      | Phe          | Leu        | Val<br>855 |              | Phe        | Gly          | Ala        | Asn<br>860 |              | Tyr        | Leu        | Met        |
| Cys<br>865      |            | ılle         | e Ser      |              |            |            |              | Gln        |              |            |            | Leu          | Pro        | Lys        | Asp<br>880 |
| Ser             | Tyr        | Leu          | ı Ile      | e Asp<br>885 |            | Phe        | e Leu        | ı Ph∈      | e Leu<br>890 |            | Arg        | Туг          | Leu        | Glu<br>895 | Val        |
| G1 <sub>2</sub> | r Pro      | Pro          | 900        |              | . Phe      | e Asp      | Thr          | Thr<br>905 |              | Gly        | Tyr        | Asn          | 910        |            | Thr        |
| Glı             | ı Alá      | a Gly<br>915 | y Met      | . Asr        | n Ala      | a Ile      | e Cys<br>920 |            | s Ser        | c Ala      | ı Gly      | 7 Cys<br>925 | s Glu      | ı Ser      | Phe        |

Ser Leu Thr Gln Lys Ile Gln Tyr Ala Ser Glu Phe Pro Asn Gln Ser 930 935 940

Tyr Val Ala Ile Ala Ala Ser Ser Trp Val Asp Asp Phe Ile Asp Trp 945 950 955 960

Leu Thr Pro Ser Ser Cys Cys Arg Ile Tyr Thr Arg Gly Pro His 965 970 975

Lys Asp Glu Phe Cys Pro Ser Thr Asp Thr Ser Phe Asn Cys Leu Lys 980 985 990

Asn Cys Met Asn Arg Thr Leu Gly Pro Val Arg Pro Thr Thr Glu Gln 995 1000 1005

Phe His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Thr Pro Asn Ile 1010 1015 1020

Arg Cys Pro Lys Gly Gly Leu Ala Ala Tyr Arg Thr Ser Val Asn 1025 1030 1035

Leu Ser Ser Asp Gly Gln Ile Ile Ala Ser Gln Phe Met Ala Tyr 1040 1045 1050

His Lys  $\,$  Pro Leu Arg Asn Ser  $\,$  Gln Asp  $\,$  Phe  $\,$  Thr Glu  $\,$  Ala Leu Arg  $\,$  1055  $\,$  1065

Ala Ser Arg Leu Leu Ala Ala As<br/>n Ile Thr Ala Glu Leu Arg Lys 1070 \$1075\$ 1080

Val Pro Gly Thr Asp Pro Asn Phe Glu Val Phe Pro Tyr Thr Ile 1085 1090 1095

Ser Asn Val Phe Tyr Gln Gln Tyr Leu Thr Val Leu Pro Glu Gly 1100 1105 1110

Ile Phe Thr Leu Ala Leu Cys Phe Val Pro Thr Phe Val Val Cys 1115 1120 1125

Tyr Leu Leu Gly Leu Asp Ile Arg Ser Gly Ile Leu Asn Leu 1130 1135 1140

Val Trp Gly Ile Ser Tyr Asn Ala Val Ser Leu Ile Asn Leu Val 1160 1165 1170

| Thr | Ala<br>1175 | Val  | Gly | Met | Ser | Val<br>1180 | Glu | Phe | Val | Ser | His<br>1185 | Ile | Thr | Arg |
|-----|-------------|------|-----|-----|-----|-------------|-----|-----|-----|-----|-------------|-----|-----|-----|
| Ser | Phe<br>1190 |      | Val | Ser | Thr | Lys<br>1195 | Pro | Thr | Arg | Leu | Glu<br>1200 | Arg | Ala | Lys |
| Asp | Ala<br>1205 |      | Ile | Phe | Met | Gly<br>1210 | Ser | Ala | Val | Phe | Ala<br>1215 | Gly | Val | Ala |
| Met | Thr<br>1220 |      | Phe | Pro | Gly | Ile<br>1225 | Leu | Ile | Leu |     | Phe<br>1230 | Ala | Gln | Ala |
| Gln | Leu<br>1235 | Ile  | Gln | Ile | Phe | Phe<br>1240 | Phe | Arg | Leu | Asn | Leu<br>1245 | Leu | Ile | Thr |
| Leu | Leu<br>1250 |      | Leu | Leu | His | Gly<br>1255 | Leu | Val | Phe | Leu | Pro<br>1260 | Val | Val | Leu |
| Ser | Туг<br>1265 |      | Gly | Pro | Asp | Val<br>1270 | Asn | Gln | Ala | Leu | Val<br>1275 | Leu | Glu | Glu |
| Lys | Leu<br>1280 |      | Thr | Glu | Ala | Ala<br>1285 | Met | Val | Ser | Glu | Pro<br>1290 | Ser | Cys | Pro |
| Gln | Tyr<br>1295 |      | Phe | Pro | Ala | Asp<br>1300 | Ala | Asn | Thr | Ser | Asp<br>1305 | Tyr | Val | Asn |
| Tyr | Gly<br>1310 |      | Asn | Pro | Glu | Phe<br>1315 | Ile | Pro | Glu | Ile | Asn<br>1320 | Ala | Ala | Ser |
| Ser | Ser<br>1325 |      | Pro | Lys | Ser | Asp<br>1330 | Gln | Lys | Phe |     |             |     |     |     |
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| <21 | 1> 3        | 999  |     |     |     |             |     |     |     |     |             |     |     |     |
| <21 | <212> DNA   |      |     |     |     |             |     |     |     |     |             |     |     |     |
| <21 | 3> M        | us s | p.  |     |     |             |     |     |     |     |             |     |     |     |
|     |             |      |     |     |     |             |     |     |     |     |             |     |     |     |

atggengeng entggeargg ntggytnytn tgggenytny tnytnaayws ngenearggn 60 garytntaya encenaenca yaargenggn ttytgyaent tytaygarga rtgyggnaar 120

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3780

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10

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155 gcc ttc tat gac gaa tgt ggg aag aac cca gag ctg tct gga agc ctc 203 Ala Phe Tyr Asp Glu Cys Gly Lys Asn Pro Glu Leu Ser Gly Ser Leu 251 atg aca ctc tcc aac gtg tcc tgc ctg tcc aac acg ccg gcc cgc aag Met Thr Leu Ser Asn Val Ser Cys Leu Ser Asn Thr Pro Ala Arg Lys atc aca ggt gat cac ctg atc cta tta cag aag atc tgc ccc cgc ctc 299 Ile Thr Gly Asp His Leu Ile Leu Leu Gln Lys Ile Cys Pro Arg Leu tac acc ggc ccc aac acc caa gcc tgc tgc tcc gcc aag cag ctg gta 347 Tyr Thr Gly Pro Asn Thr Gln Ala Cys Cys Ser Ala Lys Gln Leu Val 90 tca ctg gaa gcg agt ctg tcg atc acc aag gcc ctc ctc acc cgc tgc 395 Ser Leu Glu Ala Ser Leu Ser Ile Thr Lys Ala Leu Leu Thr Arg Cys 105 cca gcc tgc tct gac aat ttt gtg aac ctg cac tgc cac aac acg tgc 443 Pro Ala Cys Ser Asp Asn Phe Val Asn Leu His Cys His Asn Thr Cys 115 120 125

|            |                   |                   |                   |                   | ctc<br>Leu<br>135 |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 491  |
|------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|------------|-------------------|-------------------|-------------------|-------------------|------------|------|
|            |                   |                   |                   |                   | cca<br>Pro        |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 539  |
|            |                   |                   |                   |                   | cag<br>Gln        |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 587  |
|            |                   |                   |                   |                   | ctg<br>Leu        |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 635  |
|            |                   |                   |                   |                   | gcc<br>Ala        |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 683  |
|            |                   |                   |                   |                   | cca<br>Pro<br>215 |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 731  |
|            |                   |                   |                   |                   | agt<br>Ser        |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 779  |
|            |                   |                   |                   |                   | caa<br>Gln        |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 827  |
|            |                   |                   |                   |                   | tgt<br>Cys        |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 875  |
|            |                   |                   |                   |                   | ggc<br>Gly        |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 923  |
|            |                   |                   |                   |                   | ttc<br>Phe<br>295 |                   |                   |                   |                   |            |                   |                   |                   |                   |            | 971  |
| cgt<br>Arg | gtg<br>Val        | gcc<br>Ala        | ccc<br>Pro        | gcc<br>Ala<br>310 | agg<br>Arg        | gac<br>Asp        | aaa<br>Lys        | agc<br>Ser        | aag<br>Lys<br>315 | atg<br>Met | gtg<br>Val        | gac<br>Asp        | ccc<br>Pro        | aag<br>Lys<br>320 | aag<br>Lys | 1019 |
| ggc<br>Gly | acc<br>Thr        | agc<br>Ser        | ctc<br>Leu<br>325 | tct<br>Ser        | gac<br>Asp        | aag<br>Lys        | ctc<br>Leu        | agc<br>Ser<br>330 | ttc<br>Phe        | tcc<br>Ser | acc<br>Thr        | cac<br>His        | acc<br>Thr<br>335 | ctc<br>Leu        | ctt<br>Leu | 1067 |
| ggc<br>Gly | cag<br>Gln        | ttc<br>Phe<br>340 | ttc<br>Phe        | cag<br>Gln        | ggc<br>Gly        | tgg<br>Trp        | ggc<br>Gly<br>345 | acg<br>Thr        | tgg<br>Trp        | gtg<br>Val | gct<br>Ala        | tcg<br>Ser<br>350 | tgg<br>Trp        | cct<br>Pro        | ctg<br>Leu | 1115 |
| acc<br>Thr | atc<br>Ile<br>355 | ttg<br>Leu        | gtg<br>Val        | cta<br>Leu        | tct<br>Ser        | gtc<br>Val<br>360 | atc<br>Ile        | ccg<br>Pro        | gtg<br>Val        | gtg<br>Val | gcc<br>Ala<br>365 | ttg<br>Leu        | gca<br>Ala        | gcg<br>Ala        | ggc<br>Gly | 1163 |

|     |     |     |     |     |     |     |     | gac<br>Asp        |     |     |     |     |     |     |     | 1211 |
|-----|-----|-----|-----|-----|-----|-----|-----|-------------------|-----|-----|-----|-----|-----|-----|-----|------|
|     |     | _   |     | _   |     | _   |     | aaa<br>Lys        | _   |     |     | -   | _   |     |     | 1259 |
|     |     |     |     |     |     |     |     | gtg<br>Val<br>410 |     |     |     |     |     |     |     | 1307 |
|     | -   |     |     |     | -   |     | _   | ctg<br>Leu        | _   |     |     | _   |     |     | _   | 1355 |
|     |     | -   | _   | _   | _   | _   | -   | ctg<br>Leu        |     | _   |     |     | _   | _   |     | 1403 |
|     | _   |     |     |     | _   | -   |     | tcg<br>Ser        |     | _   | _   | _   | _   |     |     | 1451 |
|     |     |     |     |     |     |     |     | ccc<br>Pro        |     |     |     |     |     |     | _   | 1499 |
|     |     | _   | _   | _   |     |     | _   | ctc<br>Leu<br>490 | _   | _   |     |     | _   |     |     | 1547 |
|     |     |     |     |     |     |     |     | aac<br>Asn        |     |     |     |     |     |     |     | 1595 |
|     |     | _   | _   |     | _   | _   |     | ttt<br>Phe        | _   |     | _   | _   |     | -   | _   | 1643 |
|     |     |     |     |     |     |     | _   | ctg<br>Leu        | -   | _   | _   | _   | _   | _   | -   | 1691 |
|     |     |     |     |     |     |     |     | ctt<br>Leu        |     |     |     |     |     |     |     | 1739 |
|     |     |     |     |     |     |     |     | ctg<br>Leu<br>570 |     |     |     |     |     |     |     | 1787 |
|     |     |     |     |     |     |     |     | ctg<br>Leu        |     |     |     |     |     |     |     | 1835 |
|     |     |     |     |     |     |     |     | gcc<br>Ala        |     |     |     |     |     |     |     | 1883 |
| atg | ttc | cag | gtc | acg | ttc | atg | gct | gag               | cgc | tct | ctg | gaa | gac | gag | atc | 1931 |

| Met<br>610 | Phe        | Gln        | Val               | Thr               | Phe<br>615 | Met        | Ala        | Glu               | Arg        | Ser<br>620 | Leu        | Glu        | Asp               | Glu        | Ile<br>625 |      |
|------------|------------|------------|-------------------|-------------------|------------|------------|------------|-------------------|------------|------------|------------|------------|-------------------|------------|------------|------|
|            |            |            |                   | gct<br>Ala<br>630 |            |            |            |                   |            |            |            |            |                   |            |            | 1979 |
| _          |            |            | _                 | tac<br>Tyr        |            |            | -          | _                 | _          |            |            |            |                   |            |            | 2027 |
|            |            |            |                   | gtg<br>Val        |            |            |            |                   |            |            |            |            |                   |            |            | 2075 |
|            |            |            |                   | gga<br>Gly        |            |            |            |                   |            |            |            |            |                   |            |            | 2123 |
|            |            |            |                   | tcc<br>Ser        |            |            |            |                   |            |            |            |            |                   |            |            | 2171 |
|            |            |            |                   | ggg<br>Gly<br>710 |            |            |            |                   |            |            |            |            |                   |            |            | 2219 |
|            |            |            |                   | cgg<br>Arg        |            |            |            |                   |            |            |            |            |                   |            |            | 2267 |
|            |            |            |                   | agg<br>Arg        |            |            |            |                   |            |            |            |            |                   |            |            | 2315 |
|            |            |            |                   | ttc<br>Phe        |            |            |            |                   |            |            |            |            |                   |            |            | 2363 |
|            | Thr        |            | _                 | ctg<br>Leu        | Thr        | Ser        | Gly        |                   | Ala        | -          | Ile        |            |                   |            |            | 2411 |
|            |            |            |                   | gcc<br>Ala<br>790 |            |            |            |                   |            |            |            |            |                   |            |            | 2459 |
| cag<br>Gln | gag<br>Glu | gcc<br>Ala | tcc<br>Ser<br>805 | cgg<br>Arg        | ttg<br>Leu | gac<br>Asp | gtc<br>Val | tgc<br>Cys<br>810 | tgc<br>Cys | tgt<br>Cys | gtc<br>Val | aag<br>Lys | ccc<br>Pro<br>815 | cag<br>Gln | gag<br>Glu | 2507 |
|            |            |            | Pro               | ggc<br>Gly        |            |            |            |                   |            |            |            |            |                   |            |            | 2555 |
|            |            | Tyr        |                   | ccc<br>Pro        |            |            |            |                   |            |            |            |            |                   |            |            | 2603 |
| _          | _          | _          |                   | ctc<br>Leu        | -          | _          |            |                   |            | _          |            |            |                   |            |            | 2651 |

| 850   | 855 | 860  | 865                           |
|---|-----|--|-------------------------------|
|   |     | ctg gcc ctg ccc aag<br>Leu Ala Leu Pro Lys<br>875    |                               |
|   |     | aac cgc tac ttc gag<br>Asn Arg Tyr Phe Glu<br>895    |                               |
|   |     | ggc tac aac ttc tcc<br>Gly Tyr Asn Phe Ser<br>910    |                               |
|   |     | gca ggc tgc aac aac<br>Ala Gly Cys Asn Asn<br>925    |                               |
|   |     | gag ttc cct gag cag<br>Glu Phe Pro Glu Gln<br>940    |                               |
| 2 2   |     | gat gac ttc att gac<br>Asp Asp Phe Ile Asp<br>955    |                               |
| _   |     | ata tct ggc ccc aat<br>Ile Ser Gly Pro Asn<br>975    |                               |
|   |     | ctg aac tgc cta aag<br>Leu Asn Cys Leu Lys<br>990    |                               |
| atg agc atc acg atg<br>Met Ser Ile Thr Met<br>995 |     |  | ag ttc cat 3083<br>In Phe His |
| aag tat ctt ccc tg<br>Lys Tyr Leu Pro Tr<br>1010  |     | ac cgg ccc aac atc a<br>sp Arg Pro Asn Ile I<br>1020 |                               |
|   |     | gc acc tct gtg aac t<br>er Thr Ser Val Asn I<br>1035 |                               |
| tca gat ggc cag gt<br>Ser Asp Gly Gln Va<br>1040  |     | tt gcc att ctg tca c<br>al Ala Ile Leu Ser I<br>1050 |                               |
| ctg gag tac agt gg<br>Leu Glu Tyr Ser Gl<br>1055  |     | ct cac tgc aac ctc t<br>la His Cys Asn Leu 5<br>1065 |                               |
| -   |     | cc tat cac aag ccc o<br>la Tyr His Lys Pro 1<br>1080 |                               |
| aac tca cag gat ta<br>Asn Ser Gln Asp Ty<br>1085  |     | tg cgg gca gct cga g<br>eu Arg Ala Ala Arg (<br>1095 |                               |

| gca<br>Ala<br>1100 |            |            |            |            | gct<br>Ala<br>1105 |     |            |            |            |                    |            |            |   | 3398 |
|--------------------|------------|------------|------------|------------|--------------------|-----|------------|------------|------------|--------------------|------------|------------|---|------|
| ccg<br>Pro<br>1115 | _          |            |            | _          | ttc<br>Phe<br>1120 | Pro |            | _          |            |                    |            |            |   | 3443 |
| gag<br>Glu<br>1130 |            |            |            |            | atc<br>Ile<br>1135 | Leu |            |            |            |                    |            |            |   | 3488 |
| ctc<br>Leu<br>1145 |            |            |            |            | acc<br>Thr<br>1150 |     |            |            |            |                    |            |            |   | 3533 |
| ctg<br>Leu<br>1160 |            |            |            |            | ggc<br>Gly<br>1165 |     |            |            |            |                    |            | gtc<br>Val | _ | 3578 |
| atc<br>Ile<br>1175 |            |            |            |            | gtc<br>Val<br>1180 |     |            |            |            |                    |            | atc<br>Ile |   | 3623 |
| tac<br>Tyr<br>1190 |            |            |            |            | ctc<br>Leu<br>1195 |     |            |            |            |                    |            |            |   | 3668 |
| tct<br>Ser<br>1205 |            |            |            |            | tcc<br>Ser<br>1210 |     |            |            |            |                    |            | atc<br>Ile |   | 3713 |
| acc<br>Thr<br>1220 |            |            |            |            | ctg<br>Leu<br>1225 |     |            |            |            |                    |            |            |   | 3758 |
| atg<br>Met<br>1235 | gga<br>Gly | agt<br>Ser | gcg<br>Ala | gtg<br>Val | ttt<br>Phe<br>1240 | Ala | ggt<br>Gly | gtg<br>Val | gcc<br>Ala | atg<br>Met<br>1245 | acc<br>Thr | ctg<br>Leu |   | 3803 |
| ggc<br>Gly<br>1250 |            |            |            |            | ggc<br>Gly<br>1255 |     |            |            |            |                    |            |            |   | 3848 |
| ttc<br>Phe<br>1265 |            |            |            |            | aac<br>Asn<br>1270 |     |            |            |            |                    |            |            |   | 3893 |
| cat<br>His<br>1280 |            |            |            |            | ctg<br>Leu<br>1285 |     |            |            |            |                    |            |            |   | 3938 |
| gac<br>Asp<br>1295 |            |            |            |            | ctg<br>Leu<br>1300 |     |            |            |            |                    |            |            |   | 3983 |
|                    |            |            |            |            | atg<br>Met<br>1315 |     |            |            |            |                    |            |            |   | 4028 |

| cga<br>Arg<br>1325 |       |      |            | gct<br>Ala | gac<br>Asp<br>1330 |      |       |      | gtc<br>Val |       |       | _          |     | _    |      | 4073 |
|--------------------|-------|------|------------|------------|--------------------|------|-------|------|------------|-------|-------|------------|-----|------|------|------|
| ggt<br>Gly<br>1340 |       |      |            |            | gct<br>Ala<br>1345 | Gly  |       |      |            |       | Phe   | ttg<br>Leu |     |      |      | 4118 |
| aat<br>Asn<br>1355 |       |      | cag<br>Gln |            | tga                | taca | gcca  | ga g | gccct      | gtct  | agg   | ctcta      | atg |      |      | 4166 |
| gccc               | tgaad | cc a | aagg       | gttat      | ggg                | gatc | ttc ( | cttg | tgact      | g co  | cctt  | gaca       | cac | gccc | tcc  | 4226 |
| tcaaa              | atcci | ta g | ggga       | ggcca      | a ttc              | ccat | gag a | actg | cctgt      | cc ac | tgga  | ggat       | ggc | ctgc | tct  | 4286 |
| tgag               | gtato | cc a | ggca       | gcaco      | c act              | gatg | gct   | cctc | tgcto      | cc ca | tagt  | gggt       | CCC | cagt | ttc  | 4346 |
| caag               | tcac  | ct a | ggcc       | ttggg      | g cag              | tgcc | tcc   | tcct | gggc       | ct gg | gtct  | ggaa       | gtt | ggca | gga  | 4406 |
| acaga              | acaca | ac t | ccat       | gttt       | g tcc              | caca | ctc a | actc | actt       | c ct  | agga  | gccc       | act | tctc | atc  | 4466 |
| caac               | tttt  | cc c | ttct       | cagti      | cct                | ctct | cga . | aagt | cttaa      | at to | tgtg  | tcag       | taa | gtct | tta  | 4526 |
| acac               | gtag  | ca g | tgtc       | cctga      | a gaa              | caca | gac . | aatg | accad      | ct ac | cctg  | ggtg       | tga | tato | aca  | 4586 |
| ggag               | gcca  | ga g | agag       | gcaaa      | a ggc              | tcag | gcc . | aaga | gccaa      | ac go | tgtg  | ggag       | gcc | ggtc | ggc  | 4646 |
| agcc               | actc  | cc t | ccag       | ggcg       | c acc              | tgca | ggt   | ctgc | catco      | ca co | gcct  | tttc       | tgg | caag | aga  | 4706 |
| aggg               | ccca  | gg a | agga       | tgct       | c tca              | taag | gcc   | cagg | aagga      | at go | ctctc | ataa       | gca | cctt | ggt  | 4766 |
| catg               | gatt  | ag c | ccct       | cctg       | g aaa              | atgg | tgt   | tggg | tttg       | gt ct | ccag  | ctcc       | aat | actt | att  | 4826 |
| aagg               | ctgt  | tg c | tgcc       | agtca      | a agg              | ccac | cca   | ggag | tctga      | aa gg | gctgg | gagc       | tct | tggg | ıgct | 4886 |
| gggc               | tggt  | cc t | ccca       | tctt       | c acc              | tcgg | gcc   | tgga | tccc       | ag go | cctca | aacc       | agc | ccaa | ccc  | 4946 |
| gagc               | tttt  | gg a | cagc       | tctc       | c aga              | agca | tga   | actg | cagt       | gg ag | gatga | agat       | cct | ggct | ctg  | 5006 |
| tgct               | gtgc  | ac a | tagg       | tgtt       | t aat              | aaac | att   | tgtt | ggca       | ga aa | aaaaa | aaaa       | aaa | aaaa | ıaaa | 5066 |
| aaaa               | aaaa  | aa a | aaaa       | aaaa       | a aaa              | aaa  |       |      |            |       |       |            |     |      |      | 5092 |

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<212> PRT

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| Asp        | Ser        | Thr<br>275 | Phe        | Tyr        | Leu        | Gly        | Gln<br>280 | Met        | Pro        | Gly        | Ser        | Leu<br>285 | Val        | Leu        | Ile        |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| Ile        | ĭle<br>290 | Leu        | Cys        | Ser        | Val        | Phe<br>295 | Ala        | Val        | Val        | Thr        | Ile<br>300 | Leu        | Leu        | Val        | Gly        |
| Phe<br>305 | Arg        | Val        | Ala        | Pro        | Ala<br>310 | Arg        | Asp        | Lys        | Ser        | Lys<br>315 | Met        | Val        | Asp        | Pro        | Lys<br>320 |
| Lys        | Gly        | Thr        | Ser        | Leu<br>325 | Ser        | Asp        | Lys        | Leu        | Ser<br>330 | Phe        | Ser        | Thr        | His        | Thr<br>335 | Leu        |
| Leu        | Gly        | Gln        | Phe<br>340 | Phe        | Gln        | Gly        | Trp        | Gly<br>345 | Thr        | Trp        | Val        | Ala        | Ser<br>350 | Trp        | Pro        |

Leu Thr Ile Leu Val Leu Ser Val Ile Pro Val Val Ala Leu Ala Ala 355 360 365

Gly Leu Val Phe Thr Glu Leu Thr Thr Asp Pro Val Glu Leu Trp Ser 370 375 380

Ala Pro Asn Ser Gln Ala Arg Ser Glu Lys Ala Phe His Asp Gln His 385 390 395 400

Phe Gly Pro Phe Phe Arg Thr Asn Gln Val Ile Leu Thr Ala Pro Asn 405 410 415

Arg Ser Ser Tyr Arg Tyr Asp Ser Leu Leu Leu Gly Pro Lys Asn Phe 420 425 430

Ser Gly Ile Leu Asp Leu Asp Leu Leu Leu Glu Leu Glu Leu Gln 435 440 445

Glu Arg Leu Arg His Leu Gln Val Trp Ser Pro Glu Ala Gln Arg Asn 450 460

Ile Ser Leu Gln Asp Ile Cys Tyr Ala Pro Leu Asn Pro Asp Asn Thr 465 470 475 480

Ser Leu Tyr Asp Cys Cys Ile Asn Ser Leu Leu Gl<br/>n Tyr Phe Gl<br/>n Asn 485 490 495

Asn Arg Thr Leu Leu Leu Leu Thr Ala Asn Gln Thr Leu Met Gly Gln 500 505 510

Thr Ser Gln Val Asp Trp Lys Asp His Phe Leu Tyr Cys Ala Asn Ala Pro Leu Thr Phe Lys Asp Gly Thr Ala Leu Ala Leu Ser Cys Met Ala Asp Tyr Gly Ala Pro Val Phe Pro Phe Leu Ala Ile Gly Gly Tyr Lys Gly Lys Asp Tyr Ser Glu Ala Glu Ala Leu Ile Met Thr Phe Ser Leu Asn Asn Tyr Pro Ala Gly Asp Pro Arg Leu Ala Gln Ala Lys Leu Trp Glu Glu Ala Phe Leu Glu Glu Met Arg Ala Phe Gln Arg Arg Met Ala Gly Met Phe Gln Val Thr Phe Met Ala Glu Arg Ser Leu Glu Asp Glu Ile Asn Arg Thr Thr Ala Glu Asp Leu Pro Ile Phe Ala Thr Ser Tyr Ile Val Ile Phe Leu Tyr Ile Ser Leu Ala Leu Gly Ser Tyr Ser Ser Trp Ser Arg Val Met Val Asp Ser Lys Ala Thr Leu Gly Leu Gly Gly Val Ala Val Val Leu Gly Ala Val Met Ala Ala Met Gly Phe Phe Ser Tyr Leu Gly Ile Arg Ser Ser Leu Val Ile Leu Gln Val Val Pro Phe Leu Val Leu Ser Val Gly Ala Asp Asn Ile Phe Ile Phe Val Leu Glu Tyr Gln Arg Leu Pro Arg Arg Pro Gly Glu Pro Arg Glu Val His Ile Gly Arg Ala Leu Gly Arg Val Ala Pro Ser Met Leu Leu Cys Ser Leu 

Ser Glu Ala Ile Cys Phe Phe Leu Gly Ala Leu Thr Pro Met Pro Ala Val Arg Thr Phe Ala Leu Thr Ser Gly Leu Ala Val Ile Leu Asp Phe Leu Leu Gln Met Ser Ala Phe Val Ala Leu Leu Ser Leu Asp Ser Lys Arg Gln Glu Ala Ser Arg Leu Asp Val Cys Cys Cys Val Lys Pro Gln Glu Leu Pro Pro Gly Gln Gly Glu Gly Leu Leu Gly Phe Phe Gln Lys Ala Tyr Ala Pro Phe Leu Leu His Trp Ile Thr Arg Gly Val Val Leu Leu Phe Leu Ala Leu Phe Gly Val Ser Leu Tyr Ser Met Cys His Ile Ser Val Gly Leu Asp Gln Glu Leu Ala Leu Pro Lys Asp Ser Tyr Leu Leu Asp Tyr Phe Leu Phe Leu Asn Arg Tyr Phe Glu Val Gly Ala Pro Val Tyr Phe Val Thr Thr Leu Gly Tyr Asn Phe Ser Ser Glu Ala Gly Met Asn Ala Ile Cys Ser Ser Ala Gly Cys Asn Asn Phe Ser Phe Thr Gln Lys Ile Gln Tyr Ala Thr Glu Phe Pro Glu Gln Ser Tyr Leu Ala Ile Pro Ala Ser Ser Trp Val Asp Asp Phe Ile Asp Trp Leu Thr Pro Ser Ser Cys Cys Arg Leu Tyr Ile Ser Gly Pro Asn Lys Asp Lys Phe Cys Pro Ser Thr Val Asn Ser Leu Asn Cys Leu Lys Asn 

- Cys Met Ser Ile Thr Met Gly Ser Val Arg Pro Ser Val Glu Gln Phe 995 1000 1005
- His Lys Tyr Leu Pro Trp Phe Leu Asn Asp Arg Pro Asn Ile Lys 1010 1015 1020
- Cys Pro Lys Gly Gly Leu Ala Ala Tyr Ser Thr Ser Val Asn Leu 1025 1030 1035
- Thr Ser Asp Gly Gln Val Leu Asp Thr Val Ala Ile Leu Ser Pro 1040 1045 1050
- Arg Leu Glu Tyr Ser Gly Thr Ile Ser Ala His Cys Asn Leu Tyr 1055 1060 1065
- Leu Leu Asp Ser Ala Ser Arg Phe Met Ala Tyr His Lys Pro Leu 1070 1075 1080
- Lys Asn Ser Gln Asp Tyr Thr Glu Ala Leu Arg Ala Ala Arg Glu 1085 1090 1095
- Leu Ala Ala Asn Ile Thr Ala Asp Leu Arg Lys Val Pro Gly Thr 1100 1105 1110
- Asp Pro Ala Phe Glu Val Phe Pro Tyr Thr Ile Thr Asn Val Phe 1115 1120 1125
- Tyr Glu Gln Tyr Leu Thr Ile Leu Pro Glu Gly Leu Phe Met Leu 1130 1135 1140
- Ser Leu Cys Leu Val Pro Thr Phe Ala Val Ser Cys Leu Leu Leu 1145 1150 1155
- Gly Leu Asp Leu Arg Ser Gly Leu Leu Asn Leu Leu Ser Ile Val 1160 1165 1170
- Met Ile Leu Val Asp Thr Val Gly Phe Met Ala Leu Trp Gly Ile 1175 1180 1185
- Ser Tyr Asn Ala Val Ser Leu Ile Asn Leu Val Ser Ala Val Gly 1190 1195 1200
- Met Ser Val Glu Phe Val Ser His Ile Thr Arg Ser Phe Ala Ile 1205 1210 1215
- Ser Thr Lys Pro Thr Trp Leu Glu Arg Ala Lys Glu Ala Thr Ile

| Ser Met Gly Ser Ala Val Phe Ala Gly Val Ala Met Thr Asn Leu<br>1235 1240 1245 |  |  |  |  |  |  |  |  |  |
|---|--|--|--|--|--|--|--|--|--|
| Pro Gly Ile Leu Val Leu Gly Leu Ala Lys Ala Gln Leu Ile Gln<br>1250 1255 1260 |  |  |  |  |  |  |  |  |  |
| Ile Phe Phe Arg Leu Asn Leu Leu Ile Thr Leu Leu Gly Leu 1265 1270 1275        |  |  |  |  |  |  |  |  |  |
| Leu His Gly Leu Val Phe Leu Pro Val Ile Leu Ser Tyr Val Gly 1280 1285 1290    |  |  |  |  |  |  |  |  |  |
| Pro Asp Val Asn Pro Ala Leu Ala Leu Glu Gln Lys Arg Ala Glu<br>1295 1300 1305 |  |  |  |  |  |  |  |  |  |
| Glu Ala Val Ala Val Met Val Ala Ser Cys Pro Asn His Pro<br>1310 1315 1320     |  |  |  |  |  |  |  |  |  |
| Ser Arg Val Ser Thr Ala Asp Asn Ile Tyr Val Asn His Ser Phe<br>1325 1330 1335 |  |  |  |  |  |  |  |  |  |
| Glu Gly Ser Ile Lys Gly Ala Gly Ala Ile Ser Asn Phe Leu Pro<br>1340 1345 1350 |  |  |  |  |  |  |  |  |  |
| Asn Asn Gly Arg Gln Phe<br>1355   |  |  |  |  |  |  |  |  |  |
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